

09/869414

- 1 -

JC03 Rec'd PCT/PTC 27 JUN 2001

SEQUENCE LISTING

<110> Beinkowski et al.

<120> ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES THEREFOR

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<151> 1999-09-23

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 aatctcattc cctgctggcc aaagtcagca gaagaagggt aagtttgcca gttgctttag 1980
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 gaa 2043

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 <212> PRT
 <213> Mus musculus

<400> 8

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|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Met | Ala | Pro | Ala | Leu | His | Trp | Leu | Leu | Leu | Trp | Val | Gly | Ser | Gly | Met |
| 1 | | | | 5 | | | | | 10 | | | | | 15 | |
| Leu | Pro | Ala | Gln | Gly | Thr | His | Leu | Gly | Ile | Arg | Leu | Pro | Leu | Arg | Ser |
| | | | 20 | | | | | 25 | | | | | 30 | | |
| Gly | Leu | Ala | Gly | Pro | Pro | Leu | Gly | Leu | Arg | Leu | Pro | Arg | Glu | Thr | Asp |
| | | 35 | | | | | 40 | | | | | 45 | | | |
| Glu | Glu | Ser | Glu | Glu | Pro | Gly | Arg | Arg | Gly | Ser | Phe | Val | Glu | Met | Val |
| | 50 | | | | | 55 | | | | | 60 | | | | |
| Asp | Asn | Leu | Arg | Gly | Lys | Ser | Gly | Gln | Gly | Tyr | Tyr | Val | Glu | Met | Thr |
| 65 | | | | | 70 | | | | | 75 | | | | | 80 |
| Val | Gly | Ser | Pro | Pro | Gln | Thr | Leu | Asn | Ile | Leu | Val | Asp | Thr | Gly | Ser |
| | | | | 85 | | | | | 90 | | | | | 95 | |
| Ser | Asn | Phe | Ala | Val | Gly | Ala | Ala | Pro | His | Pro | Phe | Leu | His | Arg | Tyr |
| | | | 100 | | | | | 105 | | | | | | 110 | |
| Tyr | Gln | Arg | Gln | Leu | Ser | Ser | Thr | Tyr | Arg | Asp | Leu | Arg | Lys | Gly | Val |
| | 115 | | | | | | 120 | | | | | | 125 | | |
| Tyr | Val | Pro | Tyr | Thr | Gln | Gly | Lys | Trp | Glu | Gly | Glu | Leu | Gly | Thr | Asp |
| | 130 | | | | | 135 | | | | | 140 | | | | |
| Leu | Val | Ser | Ile | Pro | His | Gly | Pro | Asn | Val | Thr | Val | Arg | Ala | Asn | Ile |
| 145 | | | | | 150 | | | | | 155 | | | | | 160 |
| Ala | Ala | Ile | Thr | Glu | Ser | Asp | Lys | Phe | Phe | Ile | Asn | Gly | Ser | Asn | Trp |
| | | | | 165 | | | | | 170 | | | | | 175 | |
| Glu | Gly | Ile | Leu | Gly | Leu | Ala | Tyr | Ala | Glu | Ile | Ala | Arg | Pro | Asp | Asp |
| | | 180 | | | | | | 185 | | | | | 190 | | |
| Ser | Leu | Glu | Pro | Phe | Phe | Asp | Ser | Leu | Val | Lys | Gln | Thr | His | Ile | Pro |
| | 195 | | | | | | 200 | | | | | 205 | | | |
| Asn | Ile | Phe | Ser | Leu | Gln | Leu | Cys | Gly | Ala | Gly | Phe | Pro | Leu | Asn | Gln |
| | 210 | | | | | 215 | | | | | 220 | | | | |
| Thr | Glu | Ala | Leu | Ala | Ser | Val | Gly | Gly | Ser | Met | Ile | Ile | Gly | Gly | Ile |
| 225 | | | | | 230 | | | | | 235 | | | | | 240 |
| Asp | His | Ser | Leu | Tyr | Thr | Gly | Ser | Leu | Trp | Tyr | Thr | Pro | Ile | Arg | Arg |
| | | | | 245 | | | | | 250 | | | | | 255 | |
| Glu | Trp | Tyr | Tyr | Glu | Val | Ile | Ile | Val | Arg | Val | Glu | Ile | Asn | Gly | Gln |
| | | | 260 | | | | | 265 | | | | | 270 | | |
| Asp | Leu | Lys | Met | Asp | Cys | Lys | Glu | Tyr | Asn | Tyr | Asp | Lys | Ser | Ile | Val |
| | 275 | | | | | | 280 | | | | | 285 | | | |
| Asp | Ser | Gly | Thr | Thr | Asn | Leu | Arg | Leu | Pro | Lys | Lys | Val | Phe | Glu | Ala |
| | 290 | | | | | 295 | | | | | | 300 | | | |

- 10 -

Ala Val Lys Ser Ile Lys Ala Ala Ser Ser Thr Glu Lys Phe Pro Asp
305 310 315 320

Gly Phe Trp Leu Gly Glu Gln Leu Val Cys Trp Gln Ala Gly Thr Thr
325 330 335

Pro Trp Asn Ile Phe Pro Val Ile Ser Leu Tyr Leu Met Gly Glu Val
340 345 350

Thr Asn Gln Ser Phe Arg Ile Thr Ile Leu Pro Gln Gln Tyr Leu Arg
355 360 365

Pro Val Glu Asp Val Ala Thr Ser Gln Asp Asp Cys Tyr Lys Phe Ala
370 375 380

Val Ser Gln Ser Ser Thr Gly Thr Val Met Gly Ala Val Ile Met Glu
385 390 395 400

Gly Phe Tyr Val Val Phe Asp Arg Ala Arg Lys Arg Ile Gly Phe Ala
405 410 415

Val Ser Ala Cys His Val His Asp Glu Phe Arg Thr Ala Ala Val Glu
420 425 430

Gly Pro Phe Val Thr Ala Asp Met Glu Asp Cys Gly Tyr Asn Ile Pro
435 440 445

Gln Thr Asp Glu Ser Thr Leu Met Thr Ile Ala Tyr Val Met Ala Ala
450 455 460

Ile Cys Ala Leu Phe Met Leu Pro Leu Cys Leu Met Val Cys Gln Trp
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Arg Cys Leu Arg Cys Leu Arg His Gln His Asp Asp Phe Ala Asp Asp
485 490 495

Ile Ser Leu Leu Lys
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<211> 2088
<212> DNA
<213> Homo sapiens

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gagaggettg aggccaagca ccgagagaga atgtcccagg tcatgagaga atgggaagag 1020

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<210> 10

<211> 695

<212> PRT

<213> Homo sapiens

<400> 10

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      20             25             30

Gln Ile Ala Met Phe Cys Gly Arg Leu Asn Met His Met Asn Val Gln
      35             40             45

Asn Gly Lys Trp Asp Ser Asp Pro Ser Gly Thr Lys Thr Cys Ile Asp
      50             55             60

Thr Lys Glu Gly Ile Leu Gln Tyr Cys Gln Glu Val Tyr Pro Glu Leu
      65             70             75             80

Gln Ile Thr Asn Val Val Glu Ala Asn Gln Pro Val Thr Ile Gln Asn
      85             90             95

Trp Cys Lys Arg Gly Arg Lys Gln Cys Lys Thr His Pro His Phe Val
      100            105            110

Ile Pro Tyr Arg Cys Leu Val Gly Glu Phe Val Ser Asp Ala Leu Leu
      115            120            125

Val Pro Asp Lys Cys Lys Phe Leu His Gln Glu Arg Met Asp Val Cys
      130            135            140

Glu Thr His Leu His Trp His Thr Val Ala Lys Glu Thr Cys Ser Glu
      145            150            155            160

Lys Ser Thr Asn Leu His Asp Tyr Gly Met Leu Leu Pro Cys Gly Ile
      165            170            175

Asp Lys Phe Arg Gly Val Glu Phe Val Cys Cys Pro Leu Ala Glu Glu
      180            185            190
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Ser Asp Asn Val Asp Ser Ala Asp Ala Glu Glu Asp Asp Ser Asp Val
195 200 205

Trp Trp Gly Gly Ala Asp Thr Asp Tyr Ala Asp Gly Ser Glu Asp Lys
210 215 220

Val Val Glu Val Ala Glu Glu Glu Val Ala Glu Val Glu Glu Glu
225 230 235 240

Glu Ala Asp Asp Asp Glu Asp Asp Glu Asp Gly Asp Glu Val Glu Glu
245 250 255

Glu Ala Glu Glu Pro Tyr Glu Glu Ala Thr Glu Arg Thr Thr Ser Ile
260 265 270

Ala Thr Thr Thr Thr Thr Thr Thr Glu Ser Val Glu Glu Val Val Arg
275 280 285

Val Pro Thr Thr Ala Ala Ser Thr Pro Asp Ala Val Asp Lys Tyr Leu
290 295 300

Glu Thr Pro Gly Asp Glu Asn Glu His Ala His Phe Gln Lys Ala Lys
305 310 315 320

Glu Arg Leu Glu Ala Lys His Arg Glu Arg Met Ser Gln Val Met Arg
325 330 335

Glu Trp Glu Glu Ala Glu Arg Gln Ala Lys Asn Leu Pro Lys Ala Asp
340 345 350

Lys Lys Ala Val Ile Gln His Phe Gln Glu Lys Val Glu Ser Leu Glu
355 360 365

Gln Glu Ala Ala Asn Glu Arg Gln Gln Leu Val Glu Thr His Met Ala
370 375 380

Arg Val Glu Ala Met Leu Asn Asp Arg Arg Arg Leu Ala Leu Glu Asn
385 390 395 400

Tyr Ile Thr Ala Leu Gln Ala Val Pro Pro Arg Pro Arg His Val Phe
405 410 415

Asn Met Leu Lys Lys Tyr Val Arg Ala Glu Gln Lys Asp Arg Gln His
420 425 430

Thr Leu Lys His Phe Glu His Val Arg Met Val Asp Pro Lys Lys Ala
435 440 445

Ala Gln Ile Arg Ser Gln Val Met Thr His Leu Arg Val Ile Tyr Glu
450 455 460

Arg Met Asn Gln Ser Leu Ser Leu Leu Tyr Asn Val Pro Ala Val Ala
465 470 475 480

Glu Glu Ile Gln Asp Glu Val Asp Glu Leu Leu Gln Lys Glu Gln Asn
485 490 495

Tyr Ser Asp Asp Val Leu Ala Asn Met Ile Ser Glu Pro Arg Ile Ser
500 505 510

Tyr Gly Asn Asp Ala Leu Met Pro Ser Leu Thr Glu Thr Lys Thr Thr
515 520 525

Val Glu Leu Leu Pro Val Asn Gly Glu Phe Ser Leu Asp Asp Leu Gln
530 535 540

Pro Trp His Ser Phe Gly Ala Asp Ser Val Pro Ala Asn Thr Glu Asn
545 550 555 560

Glu Val Glu Pro Val Asp Ala Arg Pro Ala Ala Asp Arg Gly Leu Thr
565 570 575

Thr Arg Pro Gly Ser Gly Leu Thr Asn Ile Lys Thr Glu Glu Ile Ser
580 585 590

Glu Val Lys Met Asp Ala Glu Phe Arg His Asp Ser Gly Tyr Glu Val
595 600 605

His His Gln Lys Leu Val Phe Phe Ala Glu Asp Val Gly Ser Asn Lys
610 615 620

Gly Ala Ile Ile Gly Leu Met Val Gly Gly Val Val Ile Ala Thr Val
625 630 635 640

Ile Val Ile Thr Leu Val Met Leu Lys Lys Lys Gln Tyr Thr Ser Ile
645 650 655

His His Gly Val Val Glu Val Asp Ala Ala Val Thr Pro Glu Glu Arg
660 665 670

His Leu Ser Lys Met Gln Gln Asn Gly Tyr Glu Asn Pro Thr Tyr Lys
675 680 685

Phe Phe Glu Gln Met Gln Asn
690 695

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<212> DNA
<213> Homo sapiens

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caggagaaaag tggaatcttt ggaacaggaa gcagccaacg agagacagca gctggtggag 1140

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gtggagggttg acgccgctgt caccacagag gagcgccacc tgtccaagat gcagcagaac 2040
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<210> 12
 <211> 695
 <212> PRT
 <213> Homo sapiens

<400> 12

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| 1 | | | | 5 | | | | | | 10 | | | | 15 | |
| Ala | Leu | Glu | Val | Pro | Thr | Asp | Gly | Asn | Ala | Gly | Leu | Leu | Ala | Glu | Pro |
| | | | 20 | | | | | 25 | | | | | 30 | | |
| Gln | Ile | Ala | Met | Phe | Cys | Gly | Arg | Leu | Asn | Met | His | Met | Asn | Val | Gln |
| | 35 | | | | | | 40 | | | | | 45 | | | |
| Asn | Gly | Lys | Trp | Asp | Ser | Asp | Pro | Ser | Gly | Thr | Lys | Thr | Cys | Ile | Asp |
| | 50 | | | | | 55 | | | | | 60 | | | | |
| Thr | Lys | Glu | Gly | Ile | Leu | Gln | Tyr | Cys | Gln | Glu | Val | Tyr | Pro | Glu | Leu |
| | 65 | | | | 70 | | | | | 75 | | | | 80 | |
| Gln | Ile | Thr | Asn | Val | Val | Glu | Ala | Asn | Gln | Pro | Val | Thr | Ile | Gln | Asn |
| | | | 85 | | | | | | 90 | | | | | 95 | |
| Trp | Cys | Lys | Arg | Gly | Arg | Lys | Gln | Cys | Lys | Thr | His | Pro | His | Phe | Val |
| | | | 100 | | | | | 105 | | | | | 110 | | |
| Ile | Pro | Tyr | Arg | Cys | Leu | Val | Gly | Glu | Phe | Val | Ser | Asp | Ala | Leu | Leu |
| | | 115 | | | | | 120 | | | | | 125 | | | |
| Val | Pro | Asp | Lys | Cys | Lys | Phe | Leu | His | Gln | Glu | Arg | Met | Asp | Val | Cys |
| | 130 | | | | | 135 | | | | | 140 | | | | |
| Glu | Thr | His | Leu | His | Trp | His | Thr | Val | Ala | Lys | Glu | Thr | Cys | Ser | Glu |
| 145 | | | | | 150 | | | | | 155 | | | | 160 | |
| Lys | Ser | Thr | Asn | Leu | His | Asp | Tyr | Gly | Met | Leu | Leu | Pro | Cys | Gly | Ile |
| | | | 165 | | | | | | 170 | | | | | 175 | |
| Asp | Lys | Phe | Arg | Gly | Val | Glu | Phe | Val | Cys | Cys | Pro | Leu | Ala | Glu | Glu |
| | | | 180 | | | | | 185 | | | | | 190 | | |
| Ser | Asp | Asn | Val | Asp | Ser | Ala | Asp | Ala | Glu | Glu | Asp | Asp | Ser | Asp | Val |
| | | 195 | | | | | 200 | | | | | 205 | | | |

Trp Trp Gly Gly Ala Asp Thr Asp Tyr Ala Asp Gly Ser Glu Asp Lys
210 215 220

Val Val Glu Val Ala Glu Glu Glu Glu Val Ala Glu Val Glu Glu Glu
225 230 235 240

Glu Ala Asp Asp Asp Glu Asp Asp Glu Asp Gly Asp Glu Val Glu Glu
245 250 255

Glu Ala Glu Glu Pro Tyr Glu Glu Ala Thr Glu Arg Thr Thr Ser Ile
260 265 270

Ala Thr Thr Thr Thr Thr Thr Thr Glu Ser Val Glu Glu Val Val Arg
275 280 285

Val Pro Thr Thr Ala Ala Ser Thr Pro Asp Ala Val Asp Lys Tyr Leu
290 295 300

Glu Thr Pro Gly Asp Glu Asn Glu His Ala His Phe Gln Lys Ala Lys
305 310 315 320

Glu Arg Leu Glu Ala Lys His Arg Glu Arg Met Ser Gln Val Met Arg
325 330 335

Glu Trp Glu Glu Ala Glu Arg Gln Ala Lys Asn Leu Pro Lys Ala Asp
340 345 350

Lys Lys Ala Val Ile Gln His Phe Gln Glu Lys Val Glu Ser Leu Glu
355 360 365

Gln Glu Ala Ala Asn Glu Arg Gln Gln Leu Val Glu Thr His Met Ala
370 375 380

Arg Val Glu Ala Met Leu Asn Asp Arg Arg Arg Leu Ala Leu Glu Asn
385 390 395 400

Tyr Ile Thr Ala Leu Gln Ala Val Pro Pro Arg Pro Arg His Val Phe
405 410 415

Asn Met Leu Lys Lys Tyr Val Arg Ala Glu Gln Lys Asp Arg Gln His
420 425 430

Thr Leu Lys His Phe Glu His Val Arg Met Val Asp Pro Lys Lys Ala
435 440 445

Ala Gln Ile Arg Ser Gln Val Met Thr His Leu Arg Val Ile Tyr Glu
450 455 460

Arg Met Asn Gln Ser Leu Ser Leu Leu Tyr Asn Val Pro Ala Val Ala
465 470 475 480

Glu Glu Ile Gln Asp Glu Val Asp Glu Leu Leu Gln Lys Glu Gln Asn
485 490 495

Tyr Ser Asp Asp Val Leu Ala Asn Met Ile Ser Glu Pro Arg Ile Ser
500 505 510

Tyr Gly Asn Asp Ala Leu Met Pro Ser Leu Thr Glu Thr Lys Thr Thr
515 520 525

Val Glu Leu Leu Pro Val Asn Gly Glu Phe Ser Leu Asp Asp Leu Gln
530 535 540

- 16 -

Pro Trp His Ser Phe Gly Ala Asp Ser Val Pro Ala Asn Thr Glu Asn
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Glu Val Glu Pro Val Asp Ala Arg Pro Ala Ala Asp Arg Gly Leu Thr
565 570 575

Thr Arg Pro Gly Ser Gly Leu Thr Asn Ile Lys Thr Glu Glu Ile Ser
580 585 590

Glu Val Asn Leu Asp Ala Glu Phe Arg His Asp Ser Gly Tyr Glu Val
595 600 605

His His Gln Lys Leu Val Phe Phe Ala Glu Asp Val Gly Ser Asn Lys
610 615 620

Gly Ala Ile Ile Gly Leu Met Val Gly Gly Val Val Ile Ala Thr Val
625 630 635 640

Ile Val Ile Thr Leu Val Met Leu Lys Lys Lys Gln Tyr Thr Ser Ile
645 650 655

His His Gly Val Val Glu Val Asp Ala Ala Val Thr Pro Glu Glu Arg
660 665 670

His Leu Ser Lys Met Gln Gln Asn Gly Tyr Glu Asn Pro Thr Tyr Lys
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Phe Phe Glu Gln Met Gln Asn
690 695

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<211> 2088
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<213> Homo sapiens

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Gln Ile Ala Met Phe Cys Gly Arg Leu Asn Met His Met Asn Val Gln
      35          40          45

Asn Gly Lys Trp Asp Ser Asp Pro Ser Gly Thr Lys Thr Cys Ile Asp
      50          55          60

Thr Lys Glu Gly Ile Leu Gln Tyr Cys Gln Glu Val Tyr Pro Glu Leu
      65          70          75          80

Gln Ile Thr Asn Val Val Glu Ala Asn Gln Pro Val Thr Ile Gln Asn
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Trp Cys Lys Arg Gly Arg Lys Gln Cys Lys Thr His Pro His Phe Val
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Ile Pro Tyr Arg Cys Leu Val Gly Glu Phe Val Ser Asp Ala Leu Leu
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Val Pro Asp Lys Cys Lys Phe Leu His Gln Glu Arg Met Asp Val Cys
      130          135          140

Glu Thr His Leu His Trp His Thr Val Ala Lys Glu Thr Cys Ser Glu
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Lys Ser Thr Asn Leu His Asp Tyr Gly Met Leu Leu Pro Cys Gly Ile
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Asp Lys Phe Arg Gly Val Glu Phe Val Cys Cys Pro Leu Ala Glu Glu
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Ser Asp Asn Val Asp Ser Ala Asp Ala Glu Glu Asp Asp Ser Asp Val
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Trp Trp Gly Gly Ala Asp Thr Asp Tyr Ala Asp Gly Ser Glu Asp Lys
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Val Val Glu Val Ala Glu Glu Glu Glu Val Ala Glu Val Glu Glu Glu
      225          230          235          240

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Glu Ala Glu Glu Pro Tyr Glu Glu Ala Thr Glu Arg Thr Thr Ser Ile
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Val Pro Thr Thr Ala Ala Ser Thr Pro Asp Ala Val Asp Lys Tyr Leu
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Glu Thr Pro Gly Asp Glu Asn Glu His Ala His Phe Gln Lys Ala Lys
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Glu Arg Leu Glu Ala Lys His Arg Glu Arg Met Ser Gln Val Met Arg
325 330 335

Glu Trp Glu Glu Ala Glu Arg Gln Ala Lys Asn Leu Pro Lys Ala Asp
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Lys Lys Ala Val Ile Gln His Phe Gln Glu Lys Val Glu Ser Leu Glu
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Gln Glu Ala Ala Asn Glu Arg Gln Gln Leu Val Glu Thr His Met Ala
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Tyr Ile Thr Ala Leu Gln Ala Val Pro Pro Arg Pro Arg His Val Phe
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Asn Met Leu Lys Lys Tyr Val Arg Ala Glu Gln Lys Asp Arg Gln His
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Ala Gln Ile Arg Ser Gln Val Met Thr His Leu Arg Val Ile Tyr Glu
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Arg Met Asn Gln Ser Leu Ser Leu Leu Tyr Asn Val Pro Ala Val Ala
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Glu Glu Ile Gln Asp Glu Val Asp Glu Leu Leu Gln Lys Glu Gln Asn
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Tyr Ser Asp Asp Val Leu Ala Asn Met Ile Ser Glu Pro Arg Ile Ser
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Tyr Gly Asn Asp Ala Leu Met Pro Ser Leu Thr Glu Thr Lys Thr Thr
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Val Glu Leu Leu Pro Val Asn Gly Glu Phe Ser Leu Asp Asp Leu Gln
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Pro Trp His Ser Phe Gly Ala Asp Ser Val Pro Ala Asn Thr Glu Asn
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Glu Val Glu Pro Val Asp Ala Arg Pro Ala Ala Asp Arg Gly Leu Thr
565 570 575

Thr Arg Pro Gly Ser Gly Leu Thr Asn Ile Lys Thr Glu Glu Ile Ser
580 585 590

Glu Val Lys Met Asp Ala Glu Phe Arg His Asp Ser Gly Tyr Glu Val
595 600 605

His His Gln Lys Leu Val Phe Phe Ala Glu Asp Val Gly Ser Asn Lys
610 615 620

Gly Ala Ile Ile Gly Leu Met Val Gly Gly Val Val Ile Ala Thr Val
625 630 635 640

Ile Phe Ile Thr Leu Val Met Leu Lys Lys Lys Gln Tyr Thr Ser Ile
645 650 655

His His Gly Val Val Glu Val Asp Ala Ala Val Thr Pro Glu Glu Arg
660 665 670

His Leu Ser Lys Met Gln Gln Asn Gly Tyr Glu Asn Pro Thr Tyr Lys
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Phe Phe Glu Gln Met Gln Asn
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ggggtagagt ttgtgtgttg cccactggct gaagaaagtg acaatgtgga ttctgctgat 600
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| Ala | Leu | Glu | Val | Pro | Thr | Asp | Gly | Asn | Ala | Gly | Leu | Leu | Ala | Glu | Pro | |
| | | | 20 | | | | | 25 | | | | | 30 | | | |
| Gln | Ile | Ala | Met | Phe | Cys | Gly | Arg | Leu | Asn | Met | His | Met | Asn | Val | Gln | |
| | | 35 | | | | | 40 | | | | | 45 | | | | |
| Asn | Gly | Lys | Trp | Asp | Ser | Asp | Pro | Ser | Gly | Thr | Lys | Thr | Cys | Ile | Asp | |
| | 50 | | | | | 55 | | | | | 60 | | | | | |
| Thr | Lys | Glu | Gly | Ile | Leu | Gln | Tyr | Cys | Gln | Glu | Val | Tyr | Pro | Glu | Leu | |
| 65 | | | | | 70 | | | | | 75 | | | | | 80 | |
| Gln | Ile | Thr | Asn | Val | Val | Glu | Ala | Asn | Gln | Pro | Val | Thr | Ile | Gln | Asn | |
| | | | | 85 | | | | | 90 | | | | | 95 | | |
| Trp | Cys | Lys | Arg | Gly | Arg | Lys | Gln | Cys | Lys | Thr | His | Pro | His | Phe | Val | |
| | | | 100 | | | | | 105 | | | | | 110 | | | |
| Ile | Pro | Tyr | Arg | Cys | Leu | Val | Gly | Glu | Phe | Val | Ser | Asp | Ala | Leu | Leu | |
| | | 115 | | | | | 120 | | | | | 125 | | | | |
| Val | Pro | Asp | Lys | Cys | Lys | Phe | Leu | His | Gln | Glu | Arg | Met | Asp | Val | Cys | |
| | 130 | | | | | 135 | | | | | 140 | | | | | |
| Glu | Thr | His | Leu | His | Trp | His | Thr | Val | Ala | Lys | Glu | Thr | Cys | Ser | Glu | |
| 145 | | | | | 150 | | | | | 155 | | | | | 160 | |
| Lys | Ser | Thr | Asn | Leu | His | Asp | Tyr | Gly | Met | Leu | Leu | Pro | Cys | Gly | Ile | |
| | | | | 165 | | | | | 170 | | | | | 175 | | |
| Asp | Lys | Phe | Arg | Gly | Val | Glu | Phe | Val | Cys | Cys | Pro | Leu | Ala | Glu | Glu | |
| | | | 180 | | | | | 185 | | | | | 190 | | | |
| Ser | Asp | Asn | Val | Asp | Ser | Ala | Asp | Ala | Glu | Glu | Asp | Asp | Ser | Asp | Val | |
| | | 195 | | | | | 200 | | | | | 205 | | | | |
| Trp | Trp | Gly | Gly | Ala | Asp | Thr | Asp | Tyr | Ala | Asp | Gly | Ser | Glu | Asp | Lys | |
| | 210 | | | | | 215 | | | | | 220 | | | | | |
| Val | Val | Glu | Val | Ala | Glu | Glu | Glu | Glu | Val | Ala | Glu | Val | Glu | Glu | Glu | |
| 225 | | | | | 230 | | | | | 235 | | | | | 240 | |
| Glu | Ala | Asp | Asp | Asp | Glu | Asp | Asp | Glu | Asp | Gly | Asp | Glu | Val | Glu | Glu | |
| | | | | 245 | | | | | 250 | | | | | 255 | | |

- 21 -

Glu Ala Glu Glu Pro Tyr Glu Glu Ala Thr Glu Arg Thr Thr Ser Ile
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Ala Thr Thr Thr Thr Thr Thr Thr Glu Ser Val Glu Glu Val Val Arg
275 280 285

Val Pro Thr Thr Ala Ala Ser Thr Pro Asp Ala Val Asp Lys Tyr Leu
290 295 300

Glu Thr Pro Gly Asp Glu Asn Glu His Ala His Phe Gln Lys Ala Lys
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Glu Arg Leu Glu Ala Lys His Arg Glu Arg Met Ser Gln Val Met Arg
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Glu Trp Glu Glu Ala Glu Arg Gln Ala Lys Asn Leu Pro Lys Ala Asp
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Lys Lys Ala Val Ile Gln His Phe Gln Glu Lys Val Glu Ser Leu Glu
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Gln Glu Ala Ala Asn Glu Arg Gln Gln Leu Val Glu Thr His Met Ala
370 375 380

Arg Val Glu Ala Met Leu Asn Asp Arg Arg Arg Leu Ala Leu Glu Asn
385 390 395 400

Tyr Ile Thr Ala Leu Gln Ala Val Pro Pro Arg Pro Arg His Val Phe
405 410 415

Asn Met Leu Lys Lys Tyr Val Arg Ala Glu Gln Lys Asp Arg Gln His
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Thr Leu Lys His Phe Glu His Val Arg Met Val Asp Pro Lys Lys Ala
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Ala Gln Ile Arg Ser Gln Val Met Thr His Leu Arg Val Ile Tyr Glu
450 455 460

Arg Met Asn Gln Ser Leu Ser Leu Leu Tyr Asn Val Pro Ala Val Ala
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Glu Glu Ile Gln Asp Glu Val Asp Glu Leu Leu Gln Lys Glu Gln Asn
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Tyr Ser Asp Asp Val Leu Ala Asn Met Ile Ser Glu Pro Arg Ile Ser
500 505 510

Tyr Gly Asn Asp Ala Leu Met Pro Ser Leu Thr Glu Thr Lys Thr Thr
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Val Glu Leu Leu Pro Val Asn Gly Glu Phe Ser Leu Asp Asp Leu Gln
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Pro Trp His Ser Phe Gly Ala Asp Ser Val Pro Ala Asn Thr Glu Asn
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Glu Val Glu Pro Val Asp Ala Arg Pro Ala Ala Asp Arg Gly Leu Thr
565 570 575

Thr Arg Pro Gly Ser Gly Leu Thr Asn Ile Lys Thr Glu Glu Ile Ser
580 585 590

Glu Val Lys Met Asp Ala Glu Phe Arg His Asp Ser Gly Tyr Glu Val
595 600 605

His His Gln Lys Leu Val Phe Phe Ala Glu Asp Val Gly Ser Asn Lys
610 615 620

Gly Ala Ile Ile Gly Leu Met Val Gly Gly Val Val Ile Ala Thr Val
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Ile Val Ile Thr Leu Val Met Leu Lys Lys Lys Gln Tyr Thr Ser Ile
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His His Gly Val Val Glu Val Asp Ala Ala Val Thr Pro Glu Glu Arg
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His Leu Ser Lys Met Gln Gln Asn Gly Tyr Glu Asn Pro Thr Tyr Lys
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Phe Phe Glu Gln Met Gln Asn Lys Lys
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 <211> 697
 <212> PRT
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<400> 18

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 35 40 45
 Asn Gly Lys Trp Asp Ser Asp Pro Ser Gly Thr Lys Thr Cys Ile Asp
 50 55 60
 Thr Lys Glu Gly Ile Leu Gln Tyr Cys Gln Glu Val Tyr Pro Glu Leu
 65 70 75 80
 Gln Ile Thr Asn Val Val Glu Ala Asn Gln Pro Val Thr Ile Gln Asn
 85 90 95
 Trp Cys Lys Arg Gly Arg Lys Gln Cys Lys Thr His Pro His Phe Val
 100 105 110
 Ile Pro Tyr Arg Cys Leu Val Gly Glu Phe Val Ser Asp Ala Leu Leu
 115 120 125
 Val Pro Asp Lys Cys Lys Phe Leu His Gln Glu Arg Met Asp Val Cys
 130 135 140
 Glu Thr His Leu His Trp His Thr Val Ala Lys Glu Thr Cys Ser Glu
 145 150 155 160
 Lys Ser Thr Asn Leu His Asp Tyr Gly Met Leu Leu Pro Cys Gly Ile
 165 170 175
 Asp Lys Phe Arg Gly Val Glu Phe Val Cys Cys Pro Leu Ala Glu Glu
 180 185 190
 Ser Asp Asn Val Asp Ser Ala Asp Ala Glu Glu Asp Asp Ser Asp Val
 195 200 205
 Trp Trp Gly Gly Ala Asp Thr Asp Tyr Ala Asp Gly Ser Glu Asp Lys
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 Val Val Glu Val Ala Glu Glu Glu Glu Val Ala Glu Val Glu Glu Glu
 225 230 235 240
 Glu Ala Asp Asp Asp Glu Asp Asp Glu Asp Gly Asp Glu Val Glu Glu
 245 250 255
 Glu Ala Glu Glu Pro Tyr Glu Glu Ala Thr Glu Arg Thr Thr Ser Ile
 260 265 270

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| Val | Pro | Thr | Thr | Ala | Ala | Ser | Thr | Pro | Asp | Ala | Val | Asp | Lys | Tyr | Leu | |
| | 290 | | | | | | 295 | | | | | 300 | | | | |
| Glu | Thr | Pro | Gly | Asp | Glu | Asn | Glu | His | Ala | His | Phe | Gln | Lys | Ala | Lys | |
| 305 | | | | | 310 | | | | | 315 | | | | | 320 | |
| Glu | Arg | Leu | Glu | Ala | Lys | His | Arg | Glu | Arg | Met | Ser | Gln | Val | Met | Arg | |
| | | | | 325 | | | | | 330 | | | | | 335 | | |
| Glu | Trp | Glu | Glu | Ala | Glu | Arg | Gln | Ala | Lys | Asn | Leu | Pro | Lys | Ala | Asp | |
| | | | 340 | | | | | 345 | | | | | 350 | | | |
| Lys | Lys | Ala | Val | Ile | Gln | His | Phe | Gln | Glu | Lys | Val | Glu | Ser | Leu | Glu | |
| | 355 | | | | | | 360 | | | | | 365 | | | | |
| Gln | Glu | Ala | Ala | Asn | Glu | Arg | Gln | Gln | Leu | Val | Glu | Thr | His | Met | Ala | |
| | 370 | | | | | 375 | | | | | 380 | | | | | |
| Arg | Val | Glu | Ala | Met | Leu | Asn | Asp | Arg | Arg | Arg | Leu | Ala | Leu | Glu | Asn | |
| 385 | | | | | 390 | | | | | 395 | | | | | 400 | |
| Tyr | Ile | Thr | Ala | Leu | Gln | Ala | Val | Pro | Pro | Arg | Pro | Arg | His | Val | Phe | |
| | | | | 405 | | | | | 410 | | | | | 415 | | |
| Asn | Met | Leu | Lys | Lys | Tyr | Val | Arg | Ala | Glu | Gln | Lys | Asp | Arg | Gln | His | |
| | | 420 | | | | | | 425 | | | | | 430 | | | |
| Thr | Leu | Lys | His | Phe | Glu | His | Val | Arg | Met | Val | Asp | Pro | Lys | Lys | Ala | |
| | 435 | | | | | | 440 | | | | | 445 | | | | |
| Ala | Gln | Ile | Arg | Ser | Gln | Val | Met | Thr | His | Leu | Arg | Val | Ile | Tyr | Glu | |
| | 450 | | | | | 455 | | | | | 460 | | | | | |
| Arg | Met | Asn | Gln | Ser | Leu | Ser | Leu | Leu | Tyr | Asn | Val | Pro | Ala | Val | Ala | |
| 465 | | | | | 470 | | | | | 475 | | | | | 480 | |
| Glu | Glu | Ile | Gln | Asp | Glu | Val | Asp | Glu | Leu | Leu | Gln | Lys | Glu | Gln | Asn | |
| | | | 485 | | | | | 490 | | | | | | 495 | | |
| Tyr | Ser | Asp | Asp | Val | Leu | Ala | Asn | Met | Ile | Ser | Glu | Pro | Arg | Ile | Ser | |
| | | 500 | | | | | | 505 | | | | | 510 | | | |
| Tyr | Gly | Asn | Asp | Ala | Leu | Met | Pro | Ser | Leu | Thr | Glu | Thr | Lys | Thr | Thr | |
| | 515 | | | | | | 520 | | | | | 525 | | | | |
| Val | Glu | Leu | Leu | Pro | Val | Asn | Gly | Glu | Phe | Ser | Leu | Asp | Asp | Leu | Gln | |
| | 530 | | | | | 535 | | | | | 540 | | | | | |
| Pro | Trp | His | Ser | Phe | Gly | Ala | Asp | Ser | Val | Pro | Ala | Asn | Thr | Glu | Asn | |
| 545 | | | | | 550 | | | | | 555 | | | | | 560 | |
| Glu | Val | Glu | Pro | Val | Asp | Ala | Arg | Pro | Ala | Ala | Asp | Arg | Gly | Leu | Thr | |
| | | | | 565 | | | | | 570 | | | | | 575 | | |
| Thr | Arg | Pro | Gly | Ser | Gly | Leu | Thr | Asn | Ile | Lys | Thr | Glu | Glu | Ile | Ser | |
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 35 40 45
 Asn Gly Lys Trp Asp Ser Asp Pro Ser Gly Thr Lys Thr Cys Ile Asp
 50 55 60
 Thr Lys Glu Gly Ile Leu Gln Tyr Cys Gln Glu Val Tyr Pro Glu Leu
 65 70 75 80
 Gln Ile Thr Asn Val Val Glu Ala Asn Gln Pro Val Thr Ile Gln Asn
 85 90 95
 Trp Cys Lys Arg Gly Arg Lys Gln Cys Lys Thr His Pro His Phe Val
 100 105 110

 Ile Pro Tyr Arg Cys Leu Val Gly Glu Phe Val Ser Asp Ala Leu Leu
 115 120 125
 Val Pro Asp Lys Cys Lys Phe Leu His Gln Glu Arg Met Asp Val Cys
 130 135 140
 Glu Thr His Leu His Trp His Thr Val Ala Lys Glu Thr Cys Ser Glu
 145 150 155 160
 Lys Ser Thr Asn Leu His Asp Tyr Gly Met Leu Leu Pro Cys Gly Ile
 165 170 175
 Asp Lys Phe Arg Gly Val Glu Phe Val Cys Cys Pro Leu Ala Glu Glu
 180 185 190
 Ser Asp Asn Val Asp Ser Ala Asp Ala Glu Glu Asp Asp Ser Asp Val
 195 200 205
 Trp Trp Gly Gly Ala Asp Thr Asp Tyr Ala Asp Gly Ser Glu Asp Lys
 210 215 220
 Val Val Glu Val Ala Glu Glu Glu Glu Val Ala Glu Val Glu Glu Glu
 225 230 235 240

 Glu Ala Asp Asp Asp Glu Asp Asp Glu Asp Gly Asp Glu Val Glu Glu
 245 250 255
 Glu Ala Glu Glu Pro Tyr Glu Glu Ala Thr Glu Arg Thr Thr Ser Ile
 260 265 270
 Ala Thr Thr Thr Thr Thr Thr Thr Glu Ser Val Glu Glu Val Val Arg
 275 280 285

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| | | | | | | | | | | | | | | | |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Val | Pro | Thr | Thr | Ala | Ala | Ser | Thr | Pro | Asp | Ala | Val | Asp | Lys | Tyr | Leu |
| 290 | | | | | | 295 | | | | | 300 | | | | |
| Glu | Thr | Pro | Gly | Asp | Glu | Asn | Glu | His | Ala | His | Phe | Gln | Lys | Ala | Lys |
| 305 | | | | | 310 | | | | | 315 | | | | | 320 |
| Glu | Arg | Leu | Glu | Ala | Lys | His | Arg | Glu | Arg | Met | Ser | Gln | Val | Met | Arg |
| | | | | 325 | | | | | 330 | | | | | 335 | |
| Glu | Trp | Glu | Glu | Ala | Glu | Arg | Gln | Ala | Lys | Asn | Leu | Pro | Lys | Ala | Asp |
| | | | 340 | | | | | 345 | | | | | 350 | | |
| Lys | Lys | Ala | Val | Ile | Gln | His | Phe | Gln | Glu | Lys | Val | Glu | Ser | Leu | Glu |
| | | 355 | | | | | 360 | | | | | 365 | | | |
| Gln | Glu | Ala | Ala | Asn | Glu | Arg | Gln | Gln | Leu | Val | Glu | Thr | His | Met | Ala |
| | 370 | | | | | 375 | | | | | 380 | | | | |
| Arg | Val | Glu | Ala | Met | Leu | Asn | Asp | Arg | Arg | Arg | Leu | Ala | Leu | Glu | Asn |
| 385 | | | | | 390 | | | | | 395 | | | | | 400 |
| Tyr | Ile | Thr | Ala | Leu | Gln | Ala | Val | Pro | Pro | Arg | Pro | Arg | His | Val | Phe |
| | | | | 405 | | | | | 410 | | | | | 415 | |
| Asn | Met | Leu | Lys | Lys | Tyr | Val | Arg | Ala | Glu | Gln | Lys | Asp | Arg | Gln | His |
| | | | 420 | | | | | 425 | | | | | 430 | | |
| Thr | Leu | Lys | His | Phe | Glu | His | Val | Arg | Met | Val | Asp | Pro | Lys | Lys | Ala |
| | | 435 | | | | | 440 | | | | | 445 | | | |
| Ala | Gln | Ile | Arg | Ser | Gln | Val | Met | Thr | His | Leu | Arg | Val | Ile | Tyr | Glu |
| | 450 | | | | | 455 | | | | | 460 | | | | |
| Arg | Met | Asn | Gln | Ser | Leu | Ser | Leu | Leu | Tyr | Asn | Val | Pro | Ala | Val | Ala |
| 465 | | | | | 470 | | | | | 475 | | | | | 480 |
| Glu | Glu | Ile | Gln | Asp | Glu | Val | Asp | Glu | Leu | Leu | Gln | Lys | Glu | Gln | Asn |
| | | | 485 | | | | | | 490 | | | | | 495 | |
| Tyr | Ser | Asp | Asp | Val | Leu | Ala | Asn | Met | Ile | Ser | Glu | Pro | Arg | Ile | Ser |
| | | | 500 | | | | | 505 | | | | | 510 | | |
| Tyr | Gly | Asn | Asp | Ala | Leu | Met | Pro | Ser | Leu | Thr | Glu | Thr | Lys | Thr | Thr |
| | | 515 | | | | | 520 | | | | | 525 | | | |
| Val | Glu | Leu | Leu | Pro | Val | Asn | Gly | Glu | Phe | Ser | Leu | Asp | Asp | Leu | Gln |
| | 530 | | | | | 535 | | | | | 540 | | | | |
| Pro | Trp | His | Ser | Phe | Gly | Ala | Asp | Ser | Val | Pro | Ala | Asn | Thr | Glu | Asn |
| 545 | | | | | 550 | | | | | 555 | | | | | 560 |
| Glu | Val | Glu | Pro | Val | Asp | Ala | Arg | Pro | Ala | Ala | Asp | Arg | Gly | Leu | Thr |
| | | | | 565 | | | | | 570 | | | | | 575 | |
| Thr | Arg | Pro | Gly | Ser | Gly | Leu | Thr | Asn | Ile | Lys | Thr | Glu | Glu | Ile | Ser |
| | | | 580 | | | | | 585 | | | | | 590 | | |
| Glu | Val | Lys | Met | Asp | Ala | Glu | Phe | Arg | His | Asp | Ser | Gly | Tyr | Glu | Val |
| | | 595 | | | | | 600 | | | | | 605 | | | |
| His | His | Gln | Lys | Leu | Val | Phe | Phe | Ala | Glu | Asp | Val | Gly | Ser | Asn | Lys |
| | 610 | | | | | 615 | | | | | 620 | | | | |

Gly Ala Ile Ile Gly Leu Met Val Gly Gly Val Val Ile Ala Thr Val
625 630 635 640

Ile Phe Ile Thr Leu Val Met Leu Lys Lys Lys Gln Tyr Thr Ser Ile
645 650 655

His His Gly Val Val Glu Val Asp Ala Ala Val Thr Pro Glu Glu Arg
660 665 670

His Leu Ser Lys Met Gln Gln Asn Gly Tyr Glu Asn Pro Thr Tyr Lys
675 680 685

Phe Phe Glu Gln Met Gln Asn Lys Lys
690 695

<210> 21
<211> 1341
<212> DNA
<213> Homo sapiens

<400> 21
atggctagca tgactgggtgg acagcaaatg ggtcgcggat ccacccagca cggcatccgg 60
ctgcccctgc gcagcggcct ggggggcgcc cccctggggc tgcggctgcc ccgggagacc 120
gacgaagagc ccgaggagcc cggccggagg ggagctttg tggagatggg ggacaacctg 180
aggggcaagt cggggcaggg ctactacgtg gagatgaccg tgggcagccc cccgcagacg 240
ctcaacatcc tgggtggatac aggcagcagt aactttgcag tgggtgctgc cccccacccc 300
ttcctgcacg gctactacca gaggcagctg tccagcacat accgggacct ccggaagggt 360
gtgtatgtgc cctacaccca gggcaagtgg gaaggggagc tgggcaccga cctggtaagc 420
atcccccatg gcccacacgt cactgtgcgt gccaacattg ctgccatcac tgaatcagac 480
aagttcttca tcaacggctc caactgggaa ggcacacctg ggctggccta tgctgagatt 540
gccaggcctg acgactccct ggagccttct tttgactctc tggtaaagca gaccacgtt 600
cccaacctct tctccctgca cctttgtggg gctggcttcc ccctcaacca gtctgaagtg 660
ctggcctctg tggaggagg catgatcatt ggaggtatcg accactcgtg gtacacaggc 720
agtctctggg atacacccat ccggcgggag tgggtattatg aggtcatcat tgtcgggtg 780
gagatcaatg gacaggatct gaaaatggac tgcaaggagt acaactatga caagagcatt 840
gtggacagtg gcaccaccaa ccttcgtttg cccaagaaag tgtttgaagc tgcagtcaaa 900
tccatcaagg cagcctctc cagcgagaag ttcctgatg gtttctggct aggagagcag 960
ctggtgtgct ggcaagcagg caccacccct tgggaacattt tcccagtcac ctactctac 1020
ctaattgggtg aggttaccaa ccagtccttc cgcacacca tccttcgca gcaataacctg 1080
cggccagtgg aagatgtggc cagctcccaa gacgactgtt acaagtttgc catctcacag 1140
tcateccagg gcactgttat gggagctgtt atcatggagg gcttctacgt tgtctttgat 1200
cgggcccga aacgaattgg ctttgctgtc agcgttgcc atgtgcacga tgagttcagg 1260
acggcagcgg tgggaaggccc ttttgctacc ttggacatgg aagactgtgg ctacaacatt 1320
ccacagacag atgagtcatg a 1341

<210> 22
<211> 446
<212> PRT
<213> Homo sapiens

<400> 22

Met Ala Ser Met Thr Gly Gly Gln Gln Met Gly Arg Gly Ser Thr Gln
1 5 10 15

His Gly Ile Arg Leu Pro Leu Arg Ser Gly Leu Gly Gly Ala Pro Leu
20 25 30

Gly Leu Arg Leu Pro Arg Glu Thr Asp Glu Glu Pro Glu Glu Pro Gly
35 40 45

Arg Arg Gly Ser Phe Val Glu Met Val Asp Asn Leu Arg Gly Lys Ser
50 55 60

Gly Gln Gly Tyr Tyr Val Glu Met Thr Val Gly Ser Pro Pro Gln Thr
65 70 75 80

Leu Asn Ile Leu Val Asp Thr Gly Ser Ser Asn Phe Ala Val Gly Ala
85 90 95

Ala Pro His Pro Phe Leu His Arg Tyr Tyr Gln Arg Gln Leu Ser Ser
100 105 110

Thr Tyr Arg Asp Leu Arg Lys Gly Val Tyr Val Pro Tyr Thr Gln Gly
115 120 125

Lys Trp Glu Gly Glu Leu Gly Thr Asp Leu Val Ser Ile Pro His Gly
130 135 140

Pro Asn Val Thr Val Arg Ala Asn Ile Ala Ala Ile Thr Glu Ser Asp
145 150 155 160

Lys Phe Phe Ile Asn Gly Ser Asn Trp Glu Gly Ile Leu Gly Leu Ala
165 170 175

Tyr Ala Glu Ile Ala Arg Pro Asp Asp Ser Leu Glu Pro Phe Phe Asp
180 185 190

Ser Leu Val Lys Gln Thr His Val Pro Asn Leu Phe Ser Leu His Leu
195 200 205

Cys Gly Ala Gly Phe Pro Leu Asn Gln Ser Glu Val Leu Ala Ser Val
210 215 220

Gly Gly Ser Met Ile Ile Gly Gly Ile Asp His Ser Leu Tyr Thr Gly
225 230 235 240

Ser Leu Trp Tyr Thr Pro Ile Arg Arg Glu Trp Tyr Tyr Glu Val Ile
245 250 255

Ile Val Arg Val Glu Ile Asn Gly Gln Asp Leu Lys Met Asp Cys Lys
260 265 270

Glu Tyr Asn Tyr Asp Lys Ser Ile Val Asp Ser Gly Thr Thr Asn Leu
275 280 285

Arg Leu Pro Lys Lys Val Phe Glu Ala Ala Val Lys Ser Ile Lys Ala
290 295 300

Ala Ser Ser Thr Glu Lys Phe Pro Asp Gly Phe Trp Leu Gly Glu Gln
305 310 315 320

Leu Val Cys Trp Gln Ala Gly Thr Thr Pro Trp Asn Ile Phe Pro Val
325 330 335

Ile Ser Leu Tyr Leu Met Gly Glu Val Thr Asn Gln Ser Phe Arg Ile
340 345 350

Thr Ile Leu Pro Gln Gln Tyr Leu Arg Pro Val Glu Asp Val Ala Thr
355 360 365

Ser Gln Asp Asp Cys Tyr Lys Phe Ala Ile Ser Gln Ser Ser Thr Gly
370 375 380

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Thr Val Met Gly Ala Val Ile Met Glu Gly Phe Tyr Val Val Phe Asp
385 390 395 400

Arg Ala Arg Lys Arg Ile Gly Phe Ala Val Ser Ala Cys His Val His
405 410 415

Asp Glu Phe Arg Thr Ala Ala Val Glu Gly Pro Phe Val Thr Leu Asp
420 425 430

Met Glu Asp Cys Gly Tyr Asn Ile Pro Gln Thr Asp Glu Ser
435 440 445

<210> 23
<211> 1380
<212> DNA
<213> Homo sapiens

<400> 23
atggctagca tgactggtgg acagcaaagt ggtcgcggtat cgatgactat ctctgactct 60
ccgcgtgaac aggacggatc caccacgac ggcacccggc tgcccctgcg cagcggcctg 120
gggggcgccc ccctgggggt ggggctgccc cgggagaccg acgaagagcc caggagagccc 180
ggccggaggg gcagctttgt ggagatgggt gacaacctga ggggcaagtc ggggcagggc 240
tactacgtgg agatgaccgt gggcagcccc ccgcagacgc tcaacatcct ggtggataca 300
ggcagcagta actttgcagt ggggtgctgcc cccacccct tctgcatcg ctactaccag 360
aggcagctgt ccagcacata ccgggacctc cgggaagggtg tgtatgtgcc ctacaccag 420
ggcaagtggg aaggggagct gggcaccgac ctggttaagca tcccccatgg cccaacgctc 480
actgtgcgtg ccaacattgc tgccatcact gaacagaca agttcttcat caacggctcc 540
aactgggaag gcacccctgg gctggcctat gctgagattg ccaggcctga cgactccctg 600
gagcctttct ttgactctct ggtaaagcag acccacgttc ccaacctctt ctccctgcac 660
ctttgtggtg ctggcttccc cctcaaccag tctgaagtgc tggcctctgt cggagggagc 720
atgatcattg gaggtatcga ccactcgctg tacacaggca gtctctggta tacacccatc 780
cggcgggagt ggtattatga ggtcatcatt gtgcgggtgg agatcaatgg acaggatctg 840
aaaatggact gcaaggagta caactatgac aagagcattg tggacagtgg caccaccaac 900
cttcgtttgc ccaagaaagt gtttgaagct gcagtcaaat ccatcaaggc agcctcctcc 960
acggagaagt tccctgatgg tttctggcta ggagagcagc tgggtgtgctg gcaagcaggc 1020
accacccctt ggaacatttt cccagtcac tcactctacc taatgggtga gggtaccaac 1080
cagtccttcc gcacaccat ccttcgcag caatacctgc ggccagtgga agatgtggcc 1140
acgtcccaag acgactgtta caagtttgcc atctcacagt catccacggg cactgttatg 1200
ggagctgtta tcatggaggg cttctacgtt gtctttgac gggcccgaaa acgaattggc 1260
tttgctgtca gcgcttgcca tgtgcacgat gagttcagga cggcagcggg ggaaggccct 1320
tttgtcacct tggacatgga agactgtggc tacaacattc cacagacaga tgagtcatga 1380

<210> 24
<211> 459
<212> PRT
<213> Homo sapiens

<400> 24
Met Ala Ser Met Thr Gly Gly Gln Gln Met Gly Arg Gly Ser Met Thr
1 5 10 15

Ile Ser Asp Ser Pro Arg Glu Gln Asp Gly Ser Thr Gln His Gly Ile
20 25 30

Arg Leu Pro Leu Arg Ser Gly Leu Gly Gly Ala Pro Leu Gly Leu Arg
35 40 45

Leu Pro Arg Glu Thr Asp Glu Glu Pro Glu Glu Pro Gly Arg Arg Gly
50 55 60

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Ser Phe Val Glu Met Val Asp Asn Leu Arg Gly Lys Ser Gly Gln Gly
65 70 75 80

Tyr Tyr Val Glu Met Thr Val Gly Ser Pro Pro Gln Thr Leu Asn Ile
85 90 95

Leu Val Asp Thr Gly Ser Ser Asn Phe Ala Val Gly Ala Ala Pro His
100 105 110

Pro Phe Leu His Arg Tyr Tyr Gln Arg Gln Leu Ser Ser Thr Tyr Arg
115 120 125

Asp Leu Arg Lys Gly Val Tyr Val Pro Tyr Thr Gln Gly Lys Trp Glu
130 135 140

Gly Glu Leu Gly Thr Asp Leu Val Ser Ile Pro His Gly Pro Asn Val
145 150 155 160

Thr Val Arg Ala Asn Ile Ala Ala Ile Thr Glu Ser Asp Lys Phe Phe
165 170 175

Ile Asn Gly Ser Asn Trp Glu Gly Ile Leu Gly Leu Ala Tyr Ala Glu
180 185 190

Ile Ala Arg Pro Asp Asp Ser Leu Glu Pro Phe Phe Asp Ser Leu Val
195 200 205

Lys Gln Thr His Val Pro Asn Leu Phe Ser Leu His Leu Cys Gly Ala
210 215 220

Gly Phe Pro Leu Asn Gln Ser Glu Val Leu Ala Ser Val Gly Gly Ser
225 230 235 240

Met Ile Ile Gly Gly Ile Asp His Ser Leu Tyr Thr Gly Ser Leu Trp
245 250 255

Tyr Thr Pro Ile Arg Arg Glu Trp Tyr Tyr Glu Val Ile Ile Val Arg
260 265 270

Val Glu Ile Asn Gly Gln Asp Leu Lys Met Asp Cys Lys Glu Tyr Asn
275 280 285

Tyr Asp Lys Ser Ile Val Asp Ser Gly Thr Thr Asn Leu Arg Leu Pro
290 295 300

Lys Lys Val Phe Glu Ala Ala Val Lys Ser Ile Lys Ala Ala Ser Ser
305 310 315 320

Thr Glu Lys Phe Pro Asp Gly Phe Trp Leu Gly Glu Gln Leu Val Cys
325 330 335

Trp Gln Ala Gly Thr Thr Pro Trp Asn Ile Phe Pro Val Ile Ser Leu
340 345 350

Tyr Leu Met Gly Glu Val Thr Asn Gln Ser Phe Arg Ile Thr Ile Leu
355 360 365

Pro Gln Gln Tyr Leu Arg Pro Val Glu Asp Val Ala Thr Ser Gln Asp
370 375 380

Asp Cys Tyr Lys Phe Ala Ile Ser Gln Ser Ser Thr Gly Thr Val Met
385 390 395 400

Gly Ala Val Ile Met Glu Gly Phe Tyr Val Val Phe Asp Arg Ala Arg
 405 410 415
 Lys Arg Ile Gly Phe Ala Val Ser Ala Cys His Val His Asp Glu Phe
 420 425 430
 Arg Thr Ala Ala Val Glu Gly Pro Phe Val Thr Leu Asp Met Glu Asp
 435 440 445
 Cys Gly Tyr Asn Ile Pro Gln Thr Asp Glu Ser
 450 455

<210> 25
 <211> 1302
 <212> DNA
 <213> Homo sapiens

<400> 25
 atgactcagc atggtattcg tctgccactg cgtagcggtc tgggtgggtgc tccactgggt 60
 ctgcgtctgc cccgggagac cgacgaagag cccgaggagc ccggccggag gggcagcttt 120
 gtggagatgg tggacaacct gaggggcaag tcggggcagg gctactacgt ggagatgacc 180
 gtgggcagcc cccgcagac gctcaacatc ctgggtggata caggcagcag taactttgca 240
 gtgggtgctg ccccccaccc ctctctgcat cgctactacc agaggcagct gtccagcaca 300
 taccgggacc tccggaagg tgtgtatgtg ccctacaccc agggcaagt ggaaggggag 360
 ctgggcaccg acctggtaag catcccccat ggccccaacg tcaactgtgcg tgccaacatt 420
 gctgccatca ctgaatcaga caagttcttc atcaacggct ccaactggga aggcattctg 480
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 ctggtaaaagc agaccacgt tcccaacctc ttctccctgc acctttgtgg tgcctggctt 600
 cccctcaacc agtctgaagt gctggcctct gtcggaggga gcatgatcat tggagggtatc 660
 gaccactcgc tgtacacagg cagtctctgg tatacaccca tccggcggga gtggtattat 720
 gaggtcatca ttgtgcgggt ggagatcaat ggacaggatc tgaaaatgga ctgcaaggag 780
 tacaactatg acaagagcat tgtggacagt ggcaccacca accttcgttt gcccaagaaa 840
 gtgtttgaag ctgcagtcaa atccatcaag gcagcctcct ccacggagaa gttccctgat 900
 ggtttctggc taggagagca gctgggtgtg tggcaagcag gcaccacccc ttggaacatt 960
 ttcccagtc tctcactcta cctaattgggt gaggttacca accagtcctt ccgcatcacc 1020
 atccttccgc agcaatacct gcggccagtg gaagatgtgg ccacgtccca agacgactgt 1080
 tacaagtgtg ccatctcaca gtcattccac ggcactgtta tgggagctgt tatcatggag 1140
 ggcttctacg ttgtctttga tcgggcccga aaacgaattg gctttgctgt cagcgcttgc 1200
 catgtgcacg atgagttcag gacggcagcg gtggaaggcc cttttgtcac cttggacatg 1260
 gaagactgtg gctacaacat tccacagaca gatgagtcac ga 1302

<210> 26
 <211> 433
 <212> PRT
 <213> Homo sapiens

<400> 26
 Met Thr Gln His Gly Ile Arg Leu Pro Leu Arg Ser Gly Leu Gly Gly
 1 5 10 15
 Ala Pro Leu Gly Leu Arg Leu Pro Arg Glu Thr Asp Glu Glu Pro Glu
 20 25 30
 Glu Pro Gly Arg Arg Gly Ser Phe Val Glu Met Val Asp Asn Leu Arg
 35 40 45
 Gly Lys Ser Gly Gln Gly Tyr Tyr Val Glu Met Thr Val Gly Ser Pro
 50 55 60
 Pro Gln Thr Leu Asn Ile Leu Val Asp Thr Gly Ser Ser Asn Phe Ala
 65 70 75 80

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Val Gly Ala Ala Pro His Pro Phe Leu His Arg Tyr Tyr Gln Arg Gln
85 90 95

Leu Ser Ser Thr Tyr Arg Asp Leu Arg Lys Gly Val Tyr Val Pro Tyr
100 105 110

Thr Gln Gly Lys Trp Glu Gly Glu Leu Gly Thr Asp Leu Val Ser Ile
115 120 125

Pro His Gly Pro Asn Val Thr Val Arg Ala Asn Ile Ala Ala Ile Thr
130 135 140

Glu Ser Asp Lys Phe Phe Ile Asn Gly Ser Asn Trp Glu Gly Ile Leu
145 150 155 160

Gly Leu Ala Tyr Ala Glu Ile Ala Arg Pro Asp Asp Ser Leu Glu Pro
165 170 175

Phe Phe Asp Ser Leu Val Lys Gln Thr His Val Pro Asn Leu Phe Ser
180 185 190

Leu His Leu Cys Gly Ala Gly Phe Pro Leu Asn Gln Ser Glu Val Leu
195 200 205

Ala Ser Val Gly Gly Ser Met Ile Ile Gly Gly Ile Asp His Ser Leu
210 215 220

Tyr Thr Gly Ser Leu Trp Tyr Thr Pro Ile Arg Arg Glu Trp Tyr Tyr
225 230 235 240

Glu Val Ile Ile Val Arg Val Glu Ile Asn Gly Gln Asp Leu Lys Met
245 250 255

Asp Cys Lys Glu Tyr Asn Tyr Asp Lys Ser Ile Val Asp Ser Gly Thr
260 265 270

Thr Asn Leu Arg Leu Pro Lys Lys Val Phe Glu Ala Ala Val Lys Ser
275 280 285

Ile Lys Ala Ala Ser Ser Thr Glu Lys Phe Pro Asp Gly Phe Trp Leu
290 295 300

Gly Glu Gln Leu Val Cys Trp Gln Ala Gly Thr Thr Pro Trp Asn Ile
305 310 315 320

Phe Pro Val Ile Ser Leu Tyr Leu Met Gly Glu Val Thr Asn Gln Ser
325 330 335

Phe Arg Ile Thr Ile Leu Pro Gln Gln Tyr Leu Arg Pro Val Glu Asp
340 345 350

Val Ala Thr Ser Gln Asp Asp Cys Tyr Lys Phe Ala Ile Ser Gln Ser
355 360 365

Ser Thr Gly Thr Val Met Gly Ala Val Ile Met Glu Gly Phe Tyr Val
370 375 380

Val Phe Asp Arg Ala Arg Lys Arg Ile Gly Phe Ala Val Ser Ala Cys
385 390 395 400

His Val His Asp Glu Phe Arg Thr Ala Ala Val Glu Gly Pro Phe Val
405 410 415

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Thr Leu Asp Met Glu Asp Cys Gly Tyr Asn Ile Pro Gln Thr Asp Glu
420 425 430

Ser

<210> 27
<211> 1278
<212> DNA
<213> Homo sapiens

<400> 27
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ccgctggact ctggtatcga aaccgacgga tcctttgtgg agatggtgga caacctgagg 120
ggcaagtcgg ggcagggcta ctacgtggag atgaccgtgg gcagcccccc gcagacgctc 180
aacatcctgg tggatacagg cagcagtaac ttgacagtgg gtgctgcccc ccaccccttc 240
ctgcatcgct actaccagag gcagctgtcc agcacatacc gggacctccg gaaggggtgtg 300
tatgtgccct acacccaggg caagtgggaa ggggagctgg gcaccgacct ggtaagcatc 360
ccccatggcc ccaacgtcac tgtgcgtgcc aacattgctg ccatcactga atcagacaag 420
ttcttcatca acggctccaa ctgggaaggc atcctggggc tggcctatgc tgagattgcc 480
aggcctgacg actccctgga gcctttcttt gactctctgg taaagcagac ccacgttccc 540
aacctcttct ccctgcacct ttgtggtgct ggcttcccc tcaaccagtc tgaagtgtgtg 600
gcctctgtcg gagggagcat gatcattgga ggtatcgacc actcgctgta cacaggcagt 660
ctctggtata caccatcccg gcgggagtggt tattatgagg tcatcattgt gcgggtggag 720
atcaatggac aggatctgaa aatggactgc aaggagtaca actatgacaa gagcattgtg 780
gacagtggca ccaccaacct tcgtttgccc aagaaagtgt ttgaagctgc agtcaaattcc 840
atcaaggcag cctcctccac ggagaagttc cctgatgggt tctggctagg agagcagctg 900
gtgtgctggc aagcaggcac cacccttgg aacattttcc cagtcatctc actctacctc 960
atgggtgagg ttaccaacca gtccctccgc atcaccatcc ttccgcagca atacctgagg 1020
ccagtggaa atgtggccac gtcccaagac gactgttaca agtttgccat ctcacagtca 1080
tccacgggca ctgttatggg agctgttacc atggagggtc tctacgttgt ctttgatcgg 1140
gcccgaaaac gaattggctt tgctgtcagc gcttgccatg tgcacgatga gttcaggacg 1200
gcagcgggtg aaggcccttt tgtcaccttg gacatggaag actgtggcta caacattcca 1260
cagacagatg agtcatga 1278

<210> 28
<211> 425
<212> PRT
<213> Homo sapiens

<400> 28
Met Ala Ser Met Thr Gly Gly Gln Gln Met Gly Arg Gly Ser Met Thr
1 5 10 15
Ile Ser Asp Ser Pro Leu Asp Ser Gly Ile Glu Thr Asp Gly Ser Phe
20 25 30
Val Glu Met Val Asp Asn Leu Arg Gly Lys Ser Gly Gln Gly Tyr Tyr
35 40 45
Val Glu Met Thr Val Gly Ser Pro Pro Gln Thr Leu Asn Ile Leu Val
50 55 60
Asp Thr Gly Ser Ser Asn Phe Ala Val Gly Ala Ala Pro His Pro Phe
65 70 75 80
Leu His Arg Tyr Tyr Gln Arg Gln Leu Ser Ser Thr Tyr Arg Asp Leu
85 90 95
Arg Lys Gly Val Tyr Val Pro Tyr Thr Gln Gly Lys Trp Glu Gly Glu
100 105 110
Leu Gly Thr Asp Leu Val Ser Ile Pro His Gly Pro Asn Val Thr Val
115 120 125

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Arg Ala Asn Ile Ala Ala Ile Thr Glu Ser Asp Lys Phe Phe Ile Asn
130 135 140

Gly Ser Asn Trp Glu Gly Ile Leu Gly Leu Ala Tyr Ala Glu Ile Ala
145 150 155 160

Arg Pro Asp Asp Ser Leu Glu Pro Phe Phe Asp Ser Leu Val Lys Gln
165 170 175

Thr His Val Pro Asn Leu Phe Ser Leu His Leu Cys Gly Ala Gly Phe
180 185 190

Pro Leu Asn Gln Ser Glu Val Leu Ala Ser Val Gly Gly Ser Met Ile
195 200 205

Ile Gly Gly Ile Asp His Ser Leu Tyr Thr Gly Ser Leu Trp Tyr Thr
210 215 220

Pro Ile Arg Arg Glu Trp Tyr Tyr Glu Val Ile Ile Val Arg Val Glu
225 230 235 240

Ile Asn Gly Gln Asp Leu Lys Met Asp Cys Lys Glu Tyr Asn Tyr Asp
245 250 255

Lys Ser Ile Val Asp Ser Gly Thr Thr Asn Leu Arg Leu Pro Lys Lys
260 265 270

Val Phe Glu Ala Ala Val Lys Ser Ile Lys Ala Ala Ser Ser Thr Glu
275 280 285

Lys Phe Pro Asp Gly Phe Trp Leu Gly Glu Gln Leu Val Cys Trp Gln
290 295 300

Ala Gly Thr Thr Pro Trp Asn Ile Phe Pro Val Ile Ser Leu Tyr Leu
305 310 315 320

Met Gly Glu Val Thr Asn Gln Ser Phe Arg Ile Thr Ile Leu Pro Gln
325 330 335

Gln Tyr Leu Arg Pro Val Glu Asp Val Ala Thr Ser Gln Asp Asp Cys
340 345 350

Tyr Lys Phe Ala Ile Ser Gln Ser Ser Thr Gly Thr Val Met Gly Ala
355 360 365

Val Ile Met Glu Gly Phe Tyr Val Val Phe Asp Arg Ala Arg Lys Arg
370 375 380

Ile Gly Phe Ala Val Ser Ala Cys His Val His Asp Glu Phe Arg Thr
385 390 395 400

Ala Ala Val Glu Gly Pro Phe Val Thr Leu Asp Met Glu Asp Cys Gly
405 410 415

Tyr Asn Ile Pro Gln Thr Asp Glu Ser
420 425

<210> 29
<211> 1362
<212> DNA
<213> Homo sapiens

<400> 29
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ctgaggctgc cccgggagac cgacgaagag cccgaggagc ccggccggag gggcagcttt 180
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gtgggcagcc ccccgagac gctcaacatc ctgggtggata caggcagcag taactttgca 300
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taccgggacc tccggaaggg tgtgtatgtg ccctacaccc agggcaagtg ggaaggggag 420
ctggggcaccg acctggtaag catcccccat ggccccaacg tcactgtgcg tgccaacatt 480
gctgccatca ctgaatcaga caagttcttc atcaacggct ccaactggga aggcattctg 540
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ggcttctacg ttgtctttga tcggggccga aaacgaattg gctttgtgtg cagcgcttgc 1260
catgtgcacg atgagttcag gacggcagcg gtggaaggcc cttttgtcac cttggacatg 1320
gaagactgtg gctacaacat tccacagaca gatgagtcat ga 1382

<210> 30
<211> 453
<212> PRT
<213> Homo sapiens

<400> 30
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Leu Pro Ala His Gly Thr Gln His Gly Ile Arg Leu Pro Leu Arg Ser
20 25 30
Gly Leu Gly Gly Ala Pro Leu Gly Leu Arg Leu Pro Arg Glu Thr Asp
35 40 45
Glu Glu Pro Glu Glu Pro Gly Arg Arg Gly Ser Phe Val Glu Met Val
50 55 60
Asp Asn Leu Arg Gly Lys Ser Gly Gln Gly Tyr Tyr Val Glu Met Thr
65 70 75 80
Val Gly Ser Pro Pro Gln Thr Leu Asn Ile Leu Val Asp Thr Gly Ser
85 90 95
Ser Asn Phe Ala Val Gly Ala Ala Pro His Pro Phe Leu His Arg Tyr
100 105 110
Tyr Gln Arg Gln Leu Ser Ser Thr Tyr Arg Asp Leu Arg Lys Gly Val
115 120 125
Tyr Val Pro Tyr Thr Gln Gly Lys Trp Glu Gly Glu Leu Gly Thr Asp
130 135 140
Leu Val Ser Ile Pro His Gly Pro Asn Val Thr Val Arg Ala Asn Ile
145 150 155 160

| | | | | | | | | | | | | | | | |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Ala | Ala | Ile | Thr | Glu | Ser | Asp | Lys | Phe | Phe | Ile | Asn | Gly | Ser | Asn | Trp |
| | | | | 165 | | | | | 170 | | | | | 175 | |
| Glu | Gly | Ile | Leu | Gly | Leu | Ala | Tyr | Ala | Glu | Ile | Ala | Arg | Pro | Asp | Asp |
| | | | 180 | | | | | 185 | | | | | 190 | | |
| Ser | Leu | Glu | Pro | Phe | Phe | Asp | Ser | Leu | Val | Lys | Gln | Thr | His | Val | Pro |
| | | 195 | | | | | 200 | | | | | 205 | | | |
| Asn | Leu | Phe | Ser | Leu | Gln | Leu | Cys | Gly | Ala | Gly | Phe | Pro | Leu | Asn | Gln |
| | 210 | | | | | 215 | | | | | 220 | | | | |
| Ser | Glu | Val | Leu | Ala | Ser | Val | Gly | Gly | Ser | Met | Ile | Ile | Gly | Gly | Ile |
| 225 | | | | | 230 | | | | | 235 | | | | | 240 |
| Asp | His | Ser | Leu | Tyr | Thr | Gly | Ser | Leu | Trp | Tyr | Thr | Pro | Ile | Arg | Arg |
| | | | | 245 | | | | | 250 | | | | | 255 | |
| Glu | Trp | Tyr | Tyr | Glu | Val | Ile | Ile | Val | Arg | Val | Glu | Ile | Asn | Gly | Gln |
| | | | 260 | | | | | 265 | | | | | 270 | | |
| Asp | Leu | Lys | Met | Asp | Cys | Lys | Glu | Tyr | Asn | Tyr | Asp | Lys | Ser | Ile | Val |
| | | 275 | | | | | 280 | | | | | 285 | | | |
| Asp | Ser | Gly | Thr | Thr | Asn | Leu | Arg | Leu | Pro | Lys | Lys | Val | Phe | Glu | Ala |
| | 290 | | | | | 295 | | | | | 300 | | | | |
| Ala | Val | Lys | Ser | Ile | Lys | Ala | Ala | Ser | Ser | Thr | Glu | Lys | Phe | Pro | Asp |
| 305 | | | | | 310 | | | | | 315 | | | | | 320 |
| Gly | Phe | Trp | Leu | Gly | Glu | Gln | Leu | Val | Cys | Trp | Gln | Ala | Gly | Thr | Thr |
| | | | 325 | | | | | | 330 | | | | | 335 | |
| Pro | Trp | Asn | Ile | Phe | Pro | Val | Ile | Ser | Leu | Tyr | Leu | Met | Gly | Glu | Val |
| | | 340 | | | | | | 345 | | | | | 350 | | |
| Thr | Asn | Gln | Ser | Phe | Arg | Ile | Thr | Ile | Leu | Pro | Gln | Gln | Tyr | Leu | Arg |
| | | 355 | | | | | 360 | | | | | 365 | | | |
| Pro | Val | Glu | Asp | Val | Ala | Thr | Ser | Gln | Asp | Asp | Cys | Tyr | Lys | Phe | Ala |
| | 370 | | | | | 375 | | | | | 380 | | | | |
| Ile | Ser | Gln | Ser | Ser | Thr | Gly | Thr | Val | Met | Gly | Ala | Val | Ile | Met | Glu |
| 385 | | | | | 390 | | | | | 395 | | | | | 400 |
| Gly | Phe | Tyr | Val | Val | Phe | Asp | Arg | Ala | Arg | Lys | Arg | Ile | Gly | Phe | Ala |
| | | | 405 | | | | | | 410 | | | | | 415 | |
| Val | Ser | Ala | Cys | His | Val | His | Asp | Glu | Phe | Arg | Thr | Ala | Ala | Val | Glu |
| | | | 420 | | | | | 425 | | | | | 430 | | |
| Gly | Pro | Phe | Val | Thr | Leu | Asp | Met | Glu | Asp | Cys | Gly | Tyr | Asn | Ile | Pro |
| | | 435 | | | | | 440 | | | | | 445 | | | |
| Gln | Thr | Asp | Glu | Ser | | | | | | | | | | | |
| | | | 450 | | | | | | | | | | | | |

<210> 31
 <211> 1380
 <212> DNA

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<213> Homo sapiens

<400> 31

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ctgcggtgct cccgggagac cgacgaagag cccgaggagc ccggccggag gggcagcttt 180
gtggagatgg tggacaacct gaggggcaag tcggggcagg gctactacgt ggagatgacc 240
gtgggcagcc ccccgagac gctcaacatc ctggtggata caggcagcag taactttgca 300
gtgggtgctg cccccaccc cttcctgcat cgctactacc agaggcagct gtccagcaca 360
taccgggacc tccggaaggg tgtgtatgtg ccctacaccc agggcaagtg ggaaggggag 420
ctgggcaccg acctggttaag catcccccat ggcccccaacg tcactgtgcg tgccaacatt 480
gctgccatca ctgaatcaga caagtctctt atcaacggct ccaactggga aggcacacctg 540
gggctggcct atgctgagat tgccaggcct gacgactccc tggagcctti ctttgactct 600
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gaccactcgc tgtacacagg cagtctctgg tatacaccca tccggcggga gtggtattat 780
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ggcttctacg ttgtctttga tcgggcccga aaacgaattg gctttgctgt cagcgcttgc 1260
catgtgcacg atgagttcag gacggcagcg gtggaaggcc cttttgtcac cttggacatg 1320
gaagactgtg gctacaacat tccacagaca gatgagtcac agcagcagca gcagcagtga 1380

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<210> 32

<211> 459

<212> PRT

<213> Homo sapiens

<400> 32

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Met Ala Gln Ala Leu Pro Trp Leu Leu Leu Trp Met Gly Ala Gly Val
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          20              25              30
Gly Leu Gly Gly Ala Pro Leu Gly Leu Arg Leu Pro Arg Glu Thr Asp
          35              40              45
Glu Glu Pro Glu Glu Pro Gly Arg Arg Gly Ser Phe Val Glu Met Val
          50              55              60
Asp Asn Leu Arg Gly Lys Ser Gly Gln Gly Tyr Tyr Val Glu Met Thr
          65              70              75              80
Val Gly Ser Pro Pro Gln Thr Leu Asn Ile Leu Val Asp Thr Gly Ser
          85              90              95
Ser Asn Phe Ala Val Gly Ala Ala Pro His Pro Phe Leu His Arg Tyr
          100             105             110
Tyr Gln Arg Gln Leu Ser Ser Thr Tyr Arg Asp Leu Arg Lys Gly Val
          115             120             125
Tyr Val Pro Tyr Thr Gln Gly Lys Trp Glu Gly Glu Leu Gly Thr Asp
          130             135             140
Leu Val Ser Ile Pro His Gly Pro Asn Val Thr Val Arg Ala Asn Ile
          145             150             155             160

```

Ala Ala Ile Thr Glu Ser Asp Lys Phe Phe Ile Asn Gly Ser Asn Trp
165 170 175

Glu Gly Ile Leu Gly Leu Ala Tyr Ala Glu Ile Ala Arg Pro Asp Asp
180 185 190

Ser Leu Glu Pro Phe Phe Asp Ser Leu Val Lys Gln Thr His Val Pro
195 200 205

Asn Leu Phe Ser Leu Gln Leu Cys Gly Ala Gly Phe Pro Leu Asn Gln
210 215 220

Ser Glu Val Leu Ala Ser Val Gly Gly Ser Met Ile Ile Gly Gly Ile
225 230 235 240

Asp His Ser Leu Tyr Thr Gly Ser Leu Trp Tyr Thr Pro Ile Arg Arg
245 250 255

Glu Trp Tyr Tyr Glu Val Ile Ile Val Arg Val Glu Ile Asn Gly Gln
260 265 270

Asp Leu Lys Met Asp Cys Lys Glu Tyr Asn Tyr Asp Lys Ser Ile Val
275 280 285

Asp Ser Gly Thr Thr Asn Leu Arg Leu Pro Lys Lys Val Phe Glu Ala
290 295 300

Ala Val Lys Ser Ile Lys Ala Ala Ser Ser Thr Glu Lys Phe Pro Asp
305 310 315 320

Gly Phe Trp Leu Gly Glu Gln Leu Val Cys Trp Gln Ala Gly Thr Thr
325 330 335

Pro Trp Asn Ile Phe Pro Val Ile Ser Leu Tyr Leu Met Gly Glu Val
340 345 350

Thr Asn Gln Ser Phe Arg Ile Thr Ile Leu Pro Gln Gln Tyr Leu Arg
355 360 365

Pro Val Glu Asp Val Ala Thr Ser Gln Asp Asp Cys Tyr Lys Phe Ala
370 375 380

Ile Ser Gln Ser Ser Thr Gly Thr Val Met Gly Ala Val Ile Met Glu
385 390 395 400

Gly Phe Tyr Val Val Phe Asp Arg Ala Arg Lys Arg Ile Gly Phe Ala
405 410 415

Val Ser Ala Cys His Val His Asp Glu Phe Arg Thr Ala Ala Val Glu
420 425 430

Gly Pro Phe Val Thr Leu Asp Met Glu Asp Cys Gly Tyr Asn Ile Pro
435 440 445

Gln Thr Asp Glu Ser His His His His His His
450 455

<210> 33

<211> 25

<212> PRT
<213> Homo sapiens

<400> 33
Ser Glu Gln Gln Arg Arg Pro Arg Asp Pro Glu Val Val Asn Asp Glu
1 5 10 15
Ser Ser Leu Val Arg His Arg Trp Lys
20 25

<210> 34
<211> 19
<212> PRT
<213> Homo sapiens

<400> 34
Ser Glu Gln Leu Arg Gln Gln His Asp Asp Phe Ala Asp Asp Ile Ser
1 5 10 15
Leu Leu Lys

<210> 35
<211> 29
<212> DNA
<213> Homo sapiens

<400> 35
gtggatccac ccagcacggc atccggctg 29

<210> 36
<211> 36
<212> DNA
<213> Homo sapiens

<400> 36
gaaagctttc atgactcatc tgtctgtgga atgttg 36

<210> 37
<211> 39
<212> DNA
<213> Homo sapiens

<400> 37
gatcgatgac tatctctgac tctccgcgtg aacaggacg 39

<210> 38
<211> 39
<212> DNA
<213> Homo sapiens

<400> 38
gatccgtcct gttcacgcgg agagtcagag atagtcac 39

<210> 39
<211> 77
<212> DNA
<213> Artificial Sequence

<220>
<223> Description of Artificial Sequence: Hu-Asp2

<400> 39
cggcatccgg ctgcccctgc gtagcggctc gggtaggtgct ccactgggtc tgcgtctgcc 60
ccgggagacc gacgaag 77

<210> 40
<211> 77
<212> DNA
<213> Artificial Sequence

<220>
<223> Description of Artificial Sequence: Hu-Asp2

<400> 40
cttcgtcggg ctcccggggc agacgcagac ccagtggagc accaccaga ccgctacgca 60
ggggcagccg gatgccg 77

<210> 41
<211> 51
<212> DNA
<213> Artificial Sequence

<220>
<223> Description of Artificial Sequence: Caspase 8
Cleavage Site

<400> 41
gatcgatgac tatctctgac tctccgctgg actctggtat cgaaaccgac g 51

<210> 42
<211> 51
<212> DNA
<213> Artificial Sequence

<220>
<223> Description of Artificial Sequence: Caspase 8
Cleavage Site

<400> 42
gatccgtcgg ttccgatacc agagtcacgc ggagagtcag agatagtcac c 51

<210> 43
<211> 32
<212> DNA
<213> Homo sapiens

<400> 43
aaggatcctt tgtggagatg gtggacaacc tg 32

<210> 44
<211> 36
<212> DNA
<213> Homo sapiens

<400> 44
gaaagctttc atgactcatc tgtctgtgga atgttg 36

<210> 45
<211> 24
<212> DNA
<213> Artificial Sequence

<220>
<223> Description of Artificial Sequence: 6-His tag

<400> 45
gatcgcatca tcaccatcac catg 24

<210> 46
<211> 24
<212> DNA
<213> Artificial Sequence

<220>
<223> Description of Artificial Sequence: 6-His tag

<400> 46
gatccatggt gatggtgatg atgc 24

<210> 47
<211> 22
<212> DNA
<213> Artificial Sequence

<220>
<223> Description of Artificial Sequence: primer

<400> 47
gactgaccac tcgaccaggt tc 22

<210> 48
<211> 51
<212> DNA
<213> Artificial Sequence

<220>
<223> Description of Artificial Sequence: primer

<400> 48
cgaattaaat tccagcacac tggctacttc ttgttctgca tctcaaagaa c 51

<210> 49
<211> 26
<212> DNA
<213> Artificial Sequence

<220>
<223> Description of Artificial Sequence: primer

<400> 49
cgaattaaat tccagcacac tggcta 26

<210> 50
<211> 1287
<212> DNA
<213> Artificial Sequence

<220>
<223> Description of Artificial Sequence: Hu-Asp2(b)
delta TM

<400> 50
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ggcaccacag acggcatccg gctgcccctg cgcagcggcc tggggggcgc cccctgggg 120
ctgcggctgc cccgggagac cgacgaagag cccgaggagc ccggccggag gggcagcttt 180
gtggagatgg tggacaacct gaggggcaag tcggggcagg gctactacgt ggagatgacc 240
gtgggcagcc ccccgagac gctcaacatc ctggtggata caggcagcag taactttgca 300
gtgggtgctg cccccaccc cttcctgcat cgctactacc agaggcagct gtccagcaca 360

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taccgggacc tccggaaggg tgtgtatgtg ccctacaccc agggcaagtg ggaaggggag 420
ctgggcaccg acctggtaag catcccccat ggccccaacg tcactgtgcg tgccaacatt 480
gctgccatca ctgaatcaga caagttcttc atcaacggct ccaactggga aggcacctcg 540
gggctggcct atgctgagat tgccaggctt tgtgggtgctg gcttccccct caaccagtct 600
gaagtgtctg cctctgtcgg agggagcatg atcattggag gtatcgacca ctgctgttac 660
acaggcagtc tctggtatac acccatccgg cgggagtggg attatgaggt catcattgtg 720
cgggtggaga tcaatggaca ggatctgaaa atggactgca aggagtacaa ctatgacaag 780
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aacattccac agacagatga gtcatga 1287
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<210> 51

<211> 428

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Hu-Asp2(b)
delta TM

<400> 51

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Met Ala Gln Ala Leu Pro Trp Leu Leu Leu Trp Met Gly Ala Gly Val
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Leu Pro Ala His Gly Thr Gln His Gly Ile Arg Leu Pro Leu Arg Ser
 20 25 30
Gly Leu Gly Gly Ala Pro Leu Gly Leu Arg Leu Pro Arg Glu Thr Asp
 35 40 45
Glu Glu Pro Glu Glu Pro Gly Arg Arg Gly Ser Phe Val Glu Met Val
 50 55 60
Asp Asn Leu Arg Gly Lys Ser Gly Gln Gly Tyr Tyr Val Glu Met Thr
 65 70 75 80
Val Gly Ser Pro Pro Gln Thr Leu Asn Ile Leu Val Asp Thr Gly Ser
 85 90 95
Ser Asn Phe Ala Val Gly Ala Ala Pro His Pro Phe Leu His Arg Tyr
 100 105 110
Tyr Gln Arg Gln Leu Ser Ser Thr Tyr Arg Asp Leu Arg Lys Gly Val
 115 120 125
Tyr Val Pro Tyr Thr Gln Gly Lys Trp Glu Gly Glu Leu Gly Thr Asp
 130 135 140
Leu Val Ser Ile Pro His Gly Pro Asn Val Thr Val Arg Ala Asn Ile
 145 150 155 160
Ala Ala Ile Thr Glu Ser Asp Lys Phe Phe Ile Asn Gly Ser Asn Trp
 165 170 175
Glu Gly Ile Leu Gly Leu Ala Tyr Ala Glu Ile Ala Arg Leu Cys Gly
 180 185 190
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Ala Gly Phe Pro Leu Asn Gln Ser Glu Val Leu Ala Ser Val Gly Gly
195 200 205

Ser Met Ile Ile Gly Gly Ile Asp His Ser Leu Tyr Thr Gly Ser Leu
210 215 220

Trp Tyr Thr Pro Ile Arg Arg Glu Trp Tyr Tyr Glu Val Ile Ile Val
225 230 235 240

Arg Val Glu Ile Asn Gly Gln Asp Leu Lys Met Asp Cys Lys Glu Tyr
245 250 255

Asn Tyr Asp Lys Ser Ile Val Asp Ser Gly Thr Thr Asn Leu Arg Leu
260 265 270

Pro Lys Lys Val Phe Glu Ala Ala Val Lys Ser Ile Lys Ala Ala Ser
275 280 285

Ser Thr Glu Lys Phe Pro Asp Gly Phe Trp Leu Gly Glu Gln Leu Val
290 295 300

Cys Trp Gln Ala Gly Thr Thr Pro Trp Asn Ile Phe Pro Val Ile Ser
305 310 315 320

Leu Tyr Leu Met Gly Glu Val Thr Asn Gln Ser Phe Arg Ile Thr Ile
325 330 335

Leu Pro Gln Gln Tyr Leu Arg Pro Val Glu Asp Val Ala Thr Ser Gln
340 345 350

Asp Asp Cys Tyr Lys Phe Ala Ile Ser Gln Ser Ser Thr Gly Thr Val
355 360 365

Met Gly Ala Val Ile Met Glu Gly Phe Tyr Val Val Phe Asp Arg Ala
370 375 380

Arg Lys Arg Ile Gly Phe Ala Val Ser Ala Cys His Val His Asp Glu
385 390 395 400

Phe Arg Thr Ala Ala Val Glu Gly Pro Phe Val Thr Leu Asp Met Glu
405 410 415

Asp Cys Gly Tyr Asn Ile Pro Gln Thr Asp Glu Ser
420 425

<210> 52

<211> 1305

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Hu-Asp2(b)
delta TM

<400> 52

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ctgcggctgc cccgggagac cgacgaagag cccgaggagc ccggccggag gggcagcttt 180
gtggagatgg tggacaacct gaggggcaag tcggggcagg gctactacgt ggagatgacc 240
gtgggcagcc cccgcagac gctcaacatc ctggtggata caggcagcag taactttgca 300
gtgggtgctg cccccaccc cttcctgcat cgctactacc agaggcagct gtccagcaca 360
taccgggacc tccggaaggg tgtgtatgtg ccctacaccc agggcaagtg ggaaggggag 420

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gggctggcct atgctgagat tgccaggctt tgtgggtgctg gcttccccct caaccagtct 600
gaagtgtctg cctctgtcgg agggagcatg atcattggag gtatcgacca ctcgctgtac 660
acaggcagtc tctggtatac acccatccgg cgggagtggg attatgaggt catcattgtg 720
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ttcaggacgg cagcgttgga aggccctttt gtcaccttgg acatggaaga ctgtgggtac 1260
aacattccac agacagatga gtcacagcag cagcagcagc agtga 1305

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<210> 53

<211> 434

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Hu-Asp2(b)
delta TM

<400> 53

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Met Ala Gln Ala Leu Pro Trp Leu Leu Leu Trp Met Gly Ala Gly Val
 1             5             10             15

Leu Pro Ala His Gly Thr Gln His Gly Ile Arg Leu Pro Leu Arg Ser
      20             25             30

Gly Leu Gly Gly Ala Pro Leu Gly Leu Arg Leu Pro Arg Glu Thr Asp
      35             40             45

Glu Glu Pro Glu Glu Pro Gly Arg Arg Gly Ser Phe Val Glu Met Val
      50             55             60

Asp Asn Leu Arg Gly Lys Ser Gly Gln Gly Tyr Tyr Val Glu Met Thr
      65             70             75             80

Val Gly Ser Pro Pro Gln Thr Leu Asn Ile Leu Val Asp Thr Gly Ser
      85             90             95

Ser Asn Phe Ala Val Gly Ala Ala Pro His Pro Phe Leu His Arg Tyr
      100            105            110

Tyr Gln Arg Gln Leu Ser Ser Thr Tyr Arg Asp Leu Arg Lys Gly Val
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Tyr Val Pro Tyr Thr Gln Gly Lys Trp Glu Gly Glu Leu Gly Thr Asp
      130            135            140

Leu Val Ser Ile Pro His Gly Pro Asn Val Thr Val Arg Ala Asn Ile
      145            150            155            160

Ala Ala Ile Thr Glu Ser Asp Lys Phe Phe Ile Asn Gly Ser Asn Trp
      165            170            175

Glu Gly Ile Leu Gly Leu Ala Tyr Ala Glu Ile Ala Arg Leu Cys Gly
      180            185            190

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Ala Gly Phe Pro Leu Asn Gln Ser Glu Val Leu Ala Ser Val Gly Gly
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Ser Met Ile Ile Gly Gly Ile Asp His Ser Leu Tyr Thr Gly Ser Leu
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Trp Tyr Thr Pro Ile Arg Arg Glu Trp Tyr Tyr Glu Val Ile Ile Val
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Arg Val Glu Ile Asn Gly Gln Asp Leu Lys Met Asp Cys Lys Glu Tyr
 245 250 255

Asn Tyr Asp Lys Ser Ile Val Asp Ser Gly Thr Thr Asn Leu Arg Leu
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Pro Lys Lys Val Phe Glu Ala Ala Val Lys Ser Ile Lys Ala Ala Ser
 275 280 285

Ser Thr Glu Lys Phe Pro Asp Gly Phe Trp Leu Gly Glu Gln Leu Val
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Cys Trp Gln Ala Gly Thr Thr Pro Trp Asn Ile Phe Pro Val Ile Ser
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Leu Tyr Leu Met Gly Glu Val Thr Asn Gln Ser Phe Arg Ile Thr Ile
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Leu Pro Gln Gln Tyr Leu Arg Pro Val Glu Asp Val Ala Thr Ser Gln
 340 345 350

Asp Asp Cys Tyr Lys Phe Ala Ile Ser Gln Ser Ser Thr Gly Thr Val
 355 360 365

Met Gly Ala Val Ile Met Glu Gly Phe Tyr Val Val Phe Asp Arg Ala
 370 375 380

Arg Lys Arg Ile Gly Phe Ala Val Ser Ala Cys His Val His Asp Glu
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Phe Arg Thr Ala Ala Val Glu Gly Pro Phe Val Thr Leu Asp Met Glu
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Asp Cys Gly Tyr Asn Ile Pro Gln Thr Asp Glu Ser His His His His
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His His

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- 47 -

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35 40 45

Asn Gly Lys Trp Asp Ser Asp Pro Ser Gly Thr Lys Thr Cys Ile Asp
50 55 60

Thr Lys Glu Gly Ile Leu Gln Tyr Cys Gln Glu Val Tyr Pro Glu Leu
65 70 75 80

Gln Ile Thr Asn Val Val Glu Ala Asn Gln Pro Val Thr Ile Gln Asn
85 90 95

Trp Cys Lys Arg Gly Arg Lys Gln Cys Lys Thr His Pro His Phe Val
100 105 110

Ile Pro Tyr Arg Cys Leu Val Gly Glu Phe Val Ser Asp Ala Leu Leu
115 120 125

| | | | | | | | | | | | | | | | | | | |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Val | Pro | Asp | Lys | Cys | Lys | Phe | Leu | His | Gln | Glu | Arg | Met | Asp | Val | Cys | 130 | 135 | 140 |
| Glu | Thr | His | Leu | His | Trp | His | Thr | Val | Ala | Lys | Glu | Thr | Cys | Ser | Glu | 145 | 150 | 155 |
| Lys | Ser | Thr | Asn | Leu | His | Asp | Tyr | Gly | Met | Leu | Leu | Pro | Cys | Gly | Ile | 165 | 170 | 175 |
| Asp | Lys | Phe | Arg | Gly | Val | Glu | Phe | Val | Cys | Cys | Pro | Leu | Ala | Glu | Glu | 180 | 185 | 190 |
| Ser | Asp | Asn | Val | Asp | Ser | Ala | Asp | Ala | Glu | Glu | Asp | Asp | Ser | Asp | Val | 195 | 200 | 205 |
| Trp | Trp | Gly | Gly | Ala | Asp | Thr | Asp | Tyr | Ala | Asp | Gly | Ser | Glu | Asp | Lys | 210 | 215 | 220 |
| Val | Val | Glu | Val | Ala | Glu | Glu | Glu | Glu | Val | Ala | Glu | Val | Glu | Glu | Glu | 225 | 230 | 235 |
| Glu | Ala | Asp | Asp | Asp | Glu | Asp | Asp | Glu | Asp | Gly | Asp | Glu | Val | Glu | Glu | 245 | 250 | 255 |
| Glu | Ala | Glu | Glu | Pro | Tyr | Glu | Glu | Ala | Thr | Glu | Arg | Thr | Thr | Ser | Ile | 260 | 265 | 270 |
| Ala | Thr | Thr | Thr | Thr | Thr | Thr | Thr | Glu | Ser | Val | Glu | Glu | Val | Val | Arg | 275 | 280 | 285 |
| Glu | Val | Cys | Ser | Glu | Gln | Ala | Glu | Thr | Gly | Pro | Cys | Arg | Ala | Met | Ile | 290 | 295 | 300 |
| Ser | Arg | Trp | Tyr | Phe | Asp | Val | Thr | Glu | Gly | Lys | Cys | Ala | Pro | Phe | Phe | 305 | 310 | 315 |
| Tyr | Gly | Gly | Cys | Gly | Gly | Asn | Arg | Asn | Asn | Phe | Asp | Thr | Glu | Glu | Tyr | 325 | 330 | 335 |
| Cys | Met | Ala | Val | Cys | Gly | Ser | Ala | Met | Ser | Gln | Ser | Leu | Leu | Lys | Thr | 340 | 345 | 350 |
| Thr | Gln | Glu | Pro | Leu | Ala | Arg | Asp | Pro | Val | Lys | Leu | Pro | Thr | Thr | Ala | 355 | 360 | 365 |
| Ala | Ser | Thr | Pro | Asp | Ala | Val | Asp | Lys | Tyr | Leu | Glu | Thr | Pro | Gly | Asp | 370 | 375 | 380 |
| Glu | Asn | Glu | His | Ala | His | Phe | Gln | Lys | Ala | Lys | Glu | Arg | Leu | Glu | Ala | 385 | 390 | 395 |
| Lys | His | Arg | Glu | Arg | Met | Ser | Gln | Val | Met | Arg | Glu | Trp | Glu | Glu | Ala | 405 | 410 | 415 |
| Glu | Arg | Gln | Ala | Lys | Asn | Leu | Pro | Lys | Ala | Asp | Lys | Lys | Ala | Val | Ile | 420 | 425 | 430 |
| Gln | His | Phe | Gln | Glu | Lys | Val | Glu | Ser | Leu | Glu | Gln | Glu | Ala | Ala | Asn | 435 | 440 | 445 |
| Glu | Arg | Gln | Gln | Leu | Val | Glu | Thr | His | Met | Ala | Arg | Val | Glu | Ala | Met | 450 | 455 | 460 |

[illegible]

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35 40 45
Asn Gly Lys Trp Asp Ser Asp Pro Ser Gly Thr Lys Thr Cys Ile Asp
50 55 60

| Thr | Lys | Glu | Gly | Ile | Leu | Gln | Tyr | Cys | Gln | Glu | Val | Tyr | Pro | Glu | Leu | 65 | 70 | 75 | 80 |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Gln | Ile | Thr | Asn | Val | Val | Glu | Ala | Asn | Gln | Pro | Val | Thr | Ile | Gln | Asn | | | | |
| Trp | Cys | Lys | Arg | Gly | Arg | Lys | Gln | Cys | Lys | Thr | His | Pro | His | Phe | Val | 85 | | 90 | 95 |
| Ile | Pro | Tyr | Arg | Cys | Leu | Val | Gly | Glu | Phe | Val | Ser | Asp | Ala | Leu | Leu | 100 | | 105 | 110 |
| Val | Pro | Asp | Lys | Cys | Lys | Phe | Leu | His | Gln | Glu | Arg | Met | Asp | Val | Cys | 115 | | 120 | 125 |
| Glu | Thr | His | Leu | His | Trp | His | Thr | Val | Ala | Lys | Glu | Thr | Cys | Ser | Glu | 130 | 135 | 140 | |
| Lys | Ser | Thr | Asn | Leu | His | Asp | Tyr | Gly | Met | Leu | Leu | Pro | Cys | Gly | Ile | 145 | 150 | 155 | 160 |
| Asp | Lys | Phe | Arg | Gly | Val | Glu | Phe | Val | Cys | Cys | Pro | Leu | Ala | Glu | Glu | 165 | | 170 | 175 |
| Ser | Asp | Asn | Val | Asp | Ser | Ala | Asp | Ala | Glu | Glu | Asp | Asp | Ser | Asp | Val | 180 | 185 | | 190 |
| Trp | Trp | Gly | Gly | Ala | Asp | Thr | Asp | Tyr | Ala | Asp | Gly | Ser | Glu | Asp | Lys | 195 | 200 | 205 | |
| Val | Val | Glu | Val | Ala | Glu | Glu | Glu | Glu | Val | Ala | Glu | Val | Glu | Glu | Glu | 210 | 215 | 220 | |
| Glu | Ala | Asp | Asp | Asp | Glu | Asp | Asp | Glu | Asp | Gly | Asp | Glu | Val | Glu | Glu | 225 | 230 | 235 | 240 |
| Glu | Ala | Glu | Glu | Pro | Tyr | Glu | Glu | Ala | Thr | Glu | Arg | Thr | Thr | Ser | Ile | 245 | 250 | 255 | |
| Ala | Thr | Thr | Thr | Thr | Thr | Thr | Thr | Glu | Ser | Val | Glu | Glu | Val | Val | Arg | 260 | 265 | 270 | |
| Glu | Val | Cys | Ser | Glu | Gln | Ala | Glu | Thr | Gly | Pro | Cys | Arg | Ala | Met | Ile | 275 | 280 | 285 | |
| Ser | Arg | Trp | Tyr | Phe | Asp | Val | Thr | Glu | Gly | Lys | Cys | Ala | Pro | Phe | Phe | 290 | 295 | 300 | |
| Tyr | Gly | Gly | Cys | Gly | Gly | Asn | Arg | Asn | Asn | Phe | Asp | Thr | Glu | Glu | Tyr | 305 | 310 | 315 | 320 |
| Cys | Met | Ala | Val | Cys | Gly | Ser | Ala | Ile | Pro | Thr | Thr | Ala | Ala | Ser | Thr | 325 | 330 | 335 | |
| Pro | Asp | Ala | Val | Asp | Lys | Tyr | Leu | Glu | Thr | Pro | Gly | Asp | Glu | Asn | Glu | 340 | 345 | 350 | |
| His | Ala | His | Phe | Gln | Lys | Ala | Lys | Glu | Arg | Leu | Glu | Ala | Lys | His | Arg | 355 | 360 | 365 | |
| Glu | Arg | Met | Ser | Gln | Val | Met | Arg | Glu | Trp | Glu | Glu | Ala | Glu | Arg | Gln | 370 | 375 | 380 | |
| | | | | | | | | | | | | | | | | 385 | 390 | 395 | 400 |

| | | | |
|---|-----|-----|-----|
| Ala Lys Asn Leu Pro Lys Ala Asp Lys Lys Ala Val Ile Gln His Phe | 405 | 410 | 415 |
| Gln Glu Lys Val Glu Ser Leu Glu Gln Glu Ala Ala Asn Glu Arg Gln | 420 | 425 | 430 |
| Gln Leu Val Glu Thr His Met Ala Arg Val Glu Ala Met Leu Asn Asp | 435 | 440 | 445 |
| Arg Arg Arg Leu Ala Leu Glu Asn Tyr Ile Thr Ala Leu Gln Ala Val | 450 | 455 | 460 |
| Pro Pro Arg Pro Arg His Val Phe Asn Met Leu Lys Lys Tyr Val Arg | 465 | 470 | 475 |
| Ala Glu Gln Lys Asp Arg Gln His Thr Leu Lys His Phe Glu His Val | 485 | 490 | 495 |
| Arg Met Val Asp Pro Lys Lys Ala Ala Gln Ile Arg Ser Gln Val Met | 500 | 505 | 510 |
| Thr His Leu Arg Val Ile Tyr Glu Arg Met Asn Gln Ser Leu Ser Leu | 515 | 520 | 525 |
| Leu Tyr Asn Val Pro Ala Val Ala Glu Glu Ile Gln Asp Glu Val Asp | 530 | 535 | 540 |
| Glu Leu Leu Gln Lys Glu Gln Asn Tyr Ser Asp Asp Val Leu Ala Asn | 545 | 550 | 555 |
| Met Ile Ser Glu Pro Arg Ile Ser Tyr Gly Asn Asp Ala Leu Met Pro | 565 | 570 | 575 |
| Ser Leu Thr Glu Thr Lys Thr Thr Val Glu Leu Leu Pro Val Asn Gly | 580 | 585 | 590 |
| Glu Phe Ser Leu Asp Asp Leu Gln Pro Trp His Ser Phe Gly Ala Asp | 595 | 600 | 605 |
| Ser Val Pro Ala Asn Thr Glu Asn Glu Val Glu Pro Val Asp Ala Arg | 610 | 615 | 620 |
| Pro Ala Ala Asp Arg Gly Leu Thr Thr Arg Pro Gly Ser Gly Leu Thr | 625 | 630 | 635 |
| Asn Ile Lys Thr Glu Glu Ile Ser Glu Val Lys Met Asp Ala Glu Phe | 645 | 650 | 655 |
| Arg His Asp Ser Gly Tyr Glu Val His His Gln Lys Leu Val Phe Phe | 660 | 665 | 670 |
| Ala Glu Asp Val Gly Ser Asn Lys Gly Ala Ile Ile Gly Leu Met Val | 675 | 680 | 685 |
| Gly Gly Val Val Ile Ala Thr Val Ile Val Ile Thr Leu Val Met Leu | 690 | 695 | 700 |
| Lys Lys Lys Gln Tyr Thr Ser Ile His His Gly Val Val Glu Val Asp | 705 | 710 | 715 |
| Ala Ala Val Thr Pro Glu Glu Arg His Leu Ser Lys Met Gln Gln Asn | 725 | 730 | 735 |

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Gly Tyr Glu Asn Pro Thr Tyr Lys Phe Phe Glu Gln Met Gln Asn
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ctgaacatgc acatgaatgt ccagaatggg aagtgggatt cagatccatc agggacccaa 180
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ggggttagagt ttgtgtgttg cccactggct gaagaaagtg acaatgtgga ttctgctgat 600
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<212> PRT
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Ala Leu Glu Val Pro Thr Asp Gly Asn Ala Gly Leu Leu Ala Glu Pro
20 25 30

| | | | | | | | | | | | | | | | |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Gln | Ile | Ala | Met | Phe | Cys | Gly | Arg | Leu | Asn | Met | His | Met | Asn | Val | Gln |
| | | 35 | | | | | 40 | | | | | 45 | | | |
| Asn | Gly | Lys | Trp | Asp | Ser | Asp | Pro | Ser | Gly | Thr | Lys | Thr | Cys | Ile | Asp |
| | 50 | | | | | 55 | | | | | 60 | | | | |
| Thr | Lys | Glu | Gly | Ile | Leu | Gln | Tyr | Cys | Gln | Glu | Val | Tyr | Pro | Glu | Leu |
| 65 | | | | | 70 | | | | | 75 | | | | | 80 |
| Gln | Ile | Thr | Asn | Val | Val | Glu | Ala | Asn | Gln | Pro | Val | Thr | Ile | Gln | Asn |
| | | | 85 | | | | | | 90 | | | | | 95 | |
| Trp | Cys | Lys | Arg | Gly | Arg | Lys | Gln | Cys | Lys | Thr | His | Pro | His | Phe | Val |
| | | | 100 | | | | | 105 | | | | | 110 | | |
| Ile | Pro | Tyr | Arg | Cys | Leu | Val | Gly | Glu | Phe | Val | Ser | Asp | Ala | Leu | Leu |
| | 115 | | | | | | 120 | | | | | 125 | | | |
| Val | Pro | Asp | Lys | Cys | Lys | Phe | Leu | His | Gln | Glu | Arg | Met | Asp | Val | Cys |
| | 130 | | | | | 135 | | | | | 140 | | | | |
| Glu | Thr | His | Leu | His | Trp | His | Thr | Val | Ala | Lys | Glu | Thr | Cys | Ser | Glu |
| 145 | | | | | 150 | | | | | 155 | | | | | 160 |
| Lys | Ser | Thr | Asn | Leu | His | Asp | Tyr | Gly | Met | Leu | Leu | Pro | Cys | Gly | Ile |
| | | | | 165 | | | | | 170 | | | | | 175 | |
| Asp | Lys | Phe | Arg | Gly | Val | Glu | Phe | Val | Cys | Cys | Pro | Leu | Ala | Glu | Glu |
| | | | 180 | | | | | 185 | | | | | 190 | | |
| Ser | Asp | Asn | Val | Asp | Ser | Ala | Asp | Ala | Glu | Glu | Asp | Asp | Ser | Asp | Val |
| | | 195 | | | | | 200 | | | | | 205 | | | |
| Trp | Trp | Gly | Gly | Ala | Asp | Thr | Asp | Tyr | Ala | Asp | Gly | Ser | Glu | Asp | Lys |
| | 210 | | | | | 215 | | | | | 220 | | | | |
| Val | Val | Glu | Val | Ala | Glu | Glu | Glu | Glu | Val | Ala | Glu | Val | Glu | Glu | Glu |
| 225 | | | | | 230 | | | | | 235 | | | | | 240 |
| Glu | Ala | Asp | Asp | Asp | Glu | Asp | Asp | Glu | Asp | Gly | Asp | Glu | Val | Glu | Glu |
| | | | | 245 | | | | | 250 | | | | | 255 | |
| Glu | Ala | Glu | Glu | Pro | Tyr | Glu | Glu | Ala | Thr | Glu | Arg | Thr | Thr | Ser | Ile |
| | | | 260 | | | | | 265 | | | | | 270 | | |
| Ala | Thr | Thr | Thr | Thr | Thr | Thr | Thr | Glu | Ser | Val | Glu | Glu | Val | Val | Arg |
| | 275 | | | | | | 280 | | | | | 285 | | | |
| Glu | Val | Cys | Ser | Glu | Gln | Ala | Glu | Thr | Gly | Pro | Cys | Arg | Ala | Met | Ile |
| | 290 | | | | | 295 | | | | | 300 | | | | |
| Ser | Arg | Trp | Tyr | Phe | Asp | Val | Thr | Glu | Gly | Lys | Cys | Ala | Pro | Phe | Phe |
| 305 | | | | | 310 | | | | | 315 | | | | | 320 |
| Tyr | Gly | Gly | Cys | Gly | Gly | Asn | Arg | Asn | Asn | Phe | Asp | Thr | Glu | Glu | Tyr |
| | | | | 325 | | | | | 330 | | | | | 335 | |
| Cys | Met | Ala | Val | Cys | Gly | Ser | Ala | Met | Ser | Gln | Ser | Leu | Leu | Lys | Thr |
| | | | 340 | | | | | 345 | | | | | 350 | | |
| Thr | Gln | Glu | Pro | Leu | Ala | Arg | Asp | Pro | Val | Lys | Leu | Pro | Thr | Thr | Ala |
| | | 355 | | | | | 360 | | | | | 365 | | | |

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Ala Ser Thr Pro Asp Ala Val Asp Lys Tyr Leu Glu Thr Pro Gly Asp
370 375 380

Glu Asn Glu His Ala His Phe Gln Lys Ala Lys Glu Arg Leu Glu Ala
385 390 395 400

Lys His Arg Glu Arg Met Ser Gln Val Met Arg Glu Trp Glu Glu Ala
405 410 415

Glu Arg Gln Ala Lys Asn Leu Pro Lys Ala Asp Lys Lys Ala Val Ile
420 425 430

Gln His Phe Gln Glu Lys Val Glu Ser Leu Glu Gln Glu Ala Ala Asn
435 440 445

Glu Arg Gln Gln Leu Val Glu Thr His Met Ala Arg Val Glu Ala Met
450 455 460

Leu Asn Asp Arg Arg Arg Leu Ala Leu Glu Asn Tyr Ile Thr Ala Leu
465 470 475 480

Gln Ala Val Pro Pro Arg Pro Arg His Val Phe Asn Met Leu Lys Lys
485 490 495

Tyr Val Arg Ala Glu Gln Lys Asp Arg Gln His Thr Leu Lys His Phe
500 505 510

Glu His Val Arg Met Val Asp Pro Lys Lys Ala Ala Gln Ile Arg Ser
515 520 525

Gln Val Met Thr His Leu Arg Val Ile Tyr Glu Arg Met Asn Gln Ser
530 535 540

Leu Ser Leu Leu Tyr Asn Val Pro Ala Val Ala Glu Glu Ile Gln Asp
545 550 555 560

Glu Val Asp Glu Leu Leu Gln Lys Glu Gln Asn Tyr Ser Asp Asp Val
565 570 575

Leu Ala Asn Met Ile Ser Glu Pro Arg Ile Ser Tyr Gly Asn Asp Ala
580 585 590

Leu Met Pro Ser Leu Thr Glu Thr Lys Thr Thr Val Glu Leu Leu Pro
595 600 605

Val Asn Gly Glu Phe Ser Leu Asp Asp Leu Gln Pro Trp His Ser Phe
610 615 620

Gly Ala Asp Ser Val Pro Ala Asn Thr Glu Asn Glu Val Glu Pro Val
625 630 635 640

Asp Ala Arg Pro Ala Ala Asp Arg Gly Leu Thr Thr Arg Pro Gly Ser
645 650 655

Gly Leu Thr Asn Ile Lys Thr Glu Glu Ile Ser Glu Val Lys Met Asp
660 665 670

Ala Glu Phe Arg His Asp Ser Gly Tyr Glu Val His His Gln Lys Leu
675 680 685

Val Phe Phe Ala Glu Asp Val Gly Ser Asn Lys Gly Ala Ile Ile Gly
690 695 700

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Leu Met Val Gly Gly Val Val Ile Ala Thr Val Ile Val Ile Thr Leu
705 710 715 720

Val Met Leu Lys Lys Lys Gln Tyr Thr Ser Ile His His Gly Val Val
725 730 735

Glu Val Asp Ala Ala Val Thr Pro Glu Glu Arg His Leu Ser Lys Met
740 745 750

Gln Gln Asn Gly Tyr Glu Asn Pro Thr Tyr Lys Phe Phe Glu Gln Met
755 760 765

Gln Asn Lys Lys
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<210> 60
<211> 2259
<212> DNA
<213> Homo sapiens

<400> 60
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ctgaacatgc acatgaatgt ccagaatggg aagtgggatt cagatccatc agggaccaa 180
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gagtttgtaa gtgatgccct tctcgttcc tacaagtga aattcttaca ccaggagagg 420
atggatgttt gcgaaactca tcttcaactg cacaccgctg ccaaagagac atgcagtga 480
aagagtacca acttgcatga ctacggcatg ttgctgccct gcggaattga caagttccga 540
ggggttagagt ttgtgtgttg cccactgggt gaagaaagtg acaatgtgga ttctgctgat 600
gcggaggagg atgactcggg tgtctggtgg ggcggagcag acacagacta tgcagatggg 660
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gagtctgtgg aagaggtggt tcgagaggtg tgctctgaac aagccgagac ggggccgtgc 900
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ttggtgatgc tgaagaagaa acagtaraca tctattcatc atggtgtggt ggaggtgac 2160
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<210> 61
 <211> 753
 <212> PRT
 <213> Homo sapiens

<400> 61

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Gln Ile Ala Met Phe Cys Gly Arg Leu Asn Met His Met Asn Val Gln
      35              40              45

Asn Gly Lys Trp Asp Ser Asp Pro Ser Gly Thr Lys Thr Cys Ile Asp
      50              55              60

Thr Lys Glu Gly Ile Leu Gln Tyr Cys Gln Glu Val Tyr Pro Glu Leu
      65              70              75              80

Gln Ile Thr Asn Val Val Glu Ala Asn Gln Pro Val Thr Ile Gln Asn
      85              90              95

Trp Cys Lys Arg Gly Arg Lys Gln Cys Lys Thr His Pro His Phe Val
      100              105              110

Ile Pro Tyr Arg Cys Leu Val Gly Glu Phe Val Ser Asp Ala Leu Leu
      115              120              125

Val Pro Asp Lys Cys Lys Phe Leu His Gln Glu Arg Met Asp Val Cys
      130              135              140

Glu Thr His Leu His Trp His Thr Val Ala Lys Glu Thr Cys Ser Glu
      145              150              155              160

Lys Ser Thr Asn Leu His Asp Tyr Gly Met Leu Leu Pro Cys Gly Ile
      165              170              175

Asp Lys Phe Arg Gly Val Glu Phe Val Cys Cys Pro Leu Ala Glu Glu
      180              185              190

Ser Asp Asn Val Asp Ser Ala Asp Ala Glu Glu Asp Asp Ser Asp Val
      195              200              205

Trp Trp Gly Gly Ala Asp Thr Asp Tyr Ala Asp Gly Ser Glu Asp Lys
      210              215              220

Val Val Glu Val Ala Glu Glu Glu Glu Val Ala Glu Val Glu Glu Glu
      225              230              235              240

Glu Ala Asp Asp Asp Glu Asp Asp Glu Asp Gly Asp Glu Val Glu Glu
      245              250              255

Glu Ala Glu Glu Pro Tyr Glu Glu Ala Thr Glu Arg Thr Thr Ser Ile
      260              265              270

Ala Thr Thr Thr Thr Thr Thr Thr Glu Ser Val Glu Glu Val Val Arg
      275              280              285

Glu Val Cys Ser Glu Gln Ala Glu Thr Gly Pro Cys Arg Ala Met Ile
      290              295              300
    
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Ser Arg Trp Tyr Phe Asp Val Thr Glu Gly Lys Cys Ala Pro Phe Phe
305 310 315 320

Tyr Gly Gly Cys Gly Gly Asn Arg Asn Asn Phe Asp Thr Glu Glu Tyr
325 330 335

Cys Met Ala Val Cys Gly Ser Ala Ile Pro Thr Thr Ala Ala Ser Thr
340 345 350

Pro Asp Ala Val Asp Lys Tyr Leu Glu Thr Pro Gly Asp Glu Asn Glu
355 360 365

His Ala His Phe Gln Lys Ala Lys Glu Arg Leu Glu Ala Lys His Arg
370 375 380

Glu Arg Met Ser Gln Val Met Arg Glu Trp Glu Glu Ala Glu Arg Gln
385 390 395 400

Ala Lys Asn Leu Pro Lys Ala Asp Lys Lys Ala Val Ile Gln His Phe
405 410 415

Gln Glu Lys Val Glu Ser Leu Glu Gln Glu Ala Ala Asn Glu Arg Gln
420 425 430

Gln Leu Val Glu Thr His Met Ala Arg Val Glu Ala Met Leu Asn Asp
435 440 445

Arg Arg Arg Leu Ala Leu Glu Asn Tyr Ile Thr Ala Leu Gln Ala Val
450 455 460

Pro Pro Arg Pro Arg His Val Phe Asn Met Leu Lys Lys Tyr Val Arg
465 470 475 480

Ala Glu Gln Lys Asp Arg Gln His Thr Leu Lys His Phe Glu His Val
485 490 495

Arg Met Val Asp Pro Lys Lys Ala Ala Gln Ile Arg Ser Gln Val Met
500 505 510

Thr His Leu Arg Val Ile Tyr Glu Arg Met Asn Gln Ser Leu Ser Leu
515 520 525

Leu Tyr Asn Val Pro Ala Val Ala Glu Glu Ile Gln Asp Glu Val Asp
530 535 540

Glu Leu Leu Gln Lys Glu Gln Asn Tyr Ser Asp Asp Val Leu Ala Asn
545 550 555 560

Met Ile Ser Glu Pro Arg Ile Ser Tyr Gly Asn Asp Ala Leu Met Pro
565 570 575

Ser Leu Thr Glu Thr Lys Thr Thr Val Glu Leu Leu Pro Val Asn Gly
580 585 590

Glu Phe Ser Leu Asp Asp Leu Gln Pro Trp His Ser Phe Gly Ala Asp
595 600 605

Ser Val Pro Ala Asn Thr Glu Asn Glu Val Glu Pro Val Asp Ala Arg
610 615 620

Pro Ala Ala Asp Arg Gly Leu Thr Thr Arg Pro Gly Ser Gly Leu Thr
625 630 635 640

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Asn Ile Lys Thr Glu Glu Ile Ser Glu Val Lys Met Asp Ala Glu Phe
645 650 655
Arg His Asp Ser Gly Tyr Glu Val His His Gln Lys Leu Val Phe Phe
660 665 670
Ala Glu Asp Val Gly Ser Asn Lys Gly Ala Ile Ile Gly Leu Met Val
675 680 685
Gly Gly Val Val Ile Ala Thr Val Ile Val Ile Thr Leu Val Met Leu
690 695 700
Lys Lys Lys Gln Tyr Thr Ser Ile His His Gly Val Val Glu Val Asp
705 710 715 720
Ala Ala Val Thr Pro Glu Glu Arg His Leu Ser Lys Met Gln Gln Asn
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Gly Tyr Glu Asn Pro Thr Tyr Lys Phe Phe Glu Gln Met Gln Asn Lys
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Lys

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<210> 66
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 <223> Description of Artificial Sequence: Peptide

<400> 66
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<210> 67
 <211> 8
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<400> 67
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<210> 68
 <211> 5
 <212> PRT
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<400> 68
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<210> 69
 <211> 6
 <212> PRT
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<220>
 <223> Description of Artificial Sequence: Peptide

<400> 69

Thr Gln His Gly Ile Arg
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<210> 70
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<212> PRT
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<400> 70

Glu Thr Asp Glu Glu Pro
1 5

<210> 71
<211> 15
<212> PRT
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<220>
<223> Description of Artificial Sequence: Peptide

<400> 71

Met Cys Ala Glu Val Lys Met Asp Ala Glu Phe Lys Asp Asn Pro
1 5 10 15

<210> 72
<211> 5
<212> PRT
<213> Artificial Sequence

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<400> 72

Asp Ala Glu Phe Arg
1 5

<210> 73
<211> 5
<212> PRT
<213> Artificial Sequence

<220>
<223> Description of Artificial Sequence: synthetic

<400> 73

Ser Glu Val Asn Leu
1 5